Page 1

```
=> fil reg; d ide 1-6
FTLE 'REGISTRY' ENTERED AT 14:49:36 ON 12 DEC 2003
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```

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 11 DEC 2003 HIGHEST RN 625827-33-0 DICTIONARY FILE UPDATES: 11 DEC 2003 HIGHEST RN 625827-33-0

TSCA INFORMATION NOW CURRENT THROUGH JULY 14, 2003

Please note that search-term pricing does apply when conducting SmartSELECT searches.

L16 ANSWER 1 OF 6 REGISTRY COPYRIGHT 2003 ACS on STN

Xantidar

STEREOSEARCH

CN

FS

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

```
308068=55=5, REGISTRY *.
* Use of this CAS Registry Number alone as a search term in other STN files may
  result in incomplete search results. For additional information, enter HELP
  RN* at an online arrow prompt (=>).
    Mucopolysaccharides, heparinoids (CA INDEX NAME)
CN
OTHER NAMES:
CN Danaparoid
     Heparinoids
CN
     Lomoparan
CN
CN
     OH 10172
     Orgaran
CN
     Unspecified
MF
     MAN, CTS
CI
SR
                  IMSDRUGNEWS, IMSPATENTS, IMSRESEARCH, USPATFULL
LC
     STN Files:
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L16 ANSWER 2 OF 6 REGISTRY COPYRIGHT 2003 ACS on STN
    (114870=03=0) REGISTRY
RN
     .alpha.-D-Glucopyranoside, methyl 0-2-deoxy-6-0-sulfo-2-(sulfoamino)-
CN
     .alpha.-D-glucopyranosyl-(1.fwdarw.4)-O-.beta.-D-glucopyranuronosyl-
     (1.fwdarw.4)-0-2-deoxy-3,6-di-O-sulfo-2-(sulfoamino)-.alpha.-D-
     glucopyranosyl-(1.fwdarw.4)-O-2-O-sulfo-.alpha.-L-idopyranuronosyl-
     (1.fwdarw.4)-2-deoxy-2-(sulfoamino)-, 6-(hydrogen sulfate), decasodium
     salt (9CI) (CA INDEX NAME)
OTHER NAMES:
CN
     Arixtra
     Fondaparin sodium
CN
     Fondaparinux sodium
CN
CN
     IC 85158
     IC 851589
CN
CN
    (Org_31540___
CN
     SR 90107A
```

DR 350014-67-4

MF C31 H53 N3 O49 S8 . 10 Na

SR CA

LC STN Files: ADISINSIGHT, ADISNEWS, BEILSTEIN*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CAPLUS, CASREACT, CIN, DIOGENES, EMBASE, IMSPATENTS, IMSRESEARCH, IPA, MRCK*, PROMT, SYNTHLINE, TOXCENTER, USAN, USPATZ, USPATFULL

(*File contains numerically searchable property data)

CRN (104993-28-4)

Absolute stereochemistry.

PAGE 1-A

●10 Na

PAGE 1-B

80 REFERENCES IN FILE CA (1907 TO DATE)

2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

81 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L16 ANSWER 3 OF 6 REGISTRY COPYRIGHT 2003 ACS on STN

RN (104993-28-4) REGISTRY

.alpha.-D-Glucopyranoside, methyl O-2-deoxy-6-O-sulfo-2-(sulfoamino).alpha.-D-glucopyranosyl-(1.fwdarw.4)-O-.beta.-D-glucopyranuronosyl(1.fwdarw.4)-O-2-deoxy-3,6-di-O-sulfo-2-(sulfoamino)-.alpha.-Dglucopyranosyl-(1.fwdarw.4)-O-2-O-sulfo-.alpha.-L-idopyranuronosyl(1.fwdarw.4)-2-deoxy-2-(sulfoamino)-,6-(hydrogen sulfate) (9CI) (CA
INDEX NAME)

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Page 3

OTHER NAMES:

Fondaparinux

CN PENTA

CN SR 90107

STEREOSEARCH FS

129051-67-8, 119329-39-4, 147827-38-1, 214767-51-8, 389064-08-8, DR

393796-46-8, 393796-99-1, 412015-07-7

C31 H53 N3 O49 S8 MF

CI

SR CA

BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS,

EMBASE, IMSDRUGNEWS, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, PHAR, PIRA,

PROMT, SYNTHLINE, TOXCENTER, USPATFULL

Absolute stereochemistry.

PAGE 1-A

PAGE 1-B

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 80 REFERENCES IN FILE CA (1907 TO DATE)
- 2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 80 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L16 ANSWER-4 OF 6 REGISTRY COPYRIGHT 2003 ACS on STN RN 37270-89-6 REGISTRY

Heparin, calcium salt (9CI) (CA INDEX NAME) CN

OTHER NAMES:

Calciparin

CN

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Page 4

```
Calciparine
CN
     Calcium heparin
CN
     Calcium heparinate
CN
     Ecasolv
CN
     Hepacarin
CN
     Heparin calcium
    Nadroparin calcium
101921-20-4, 39363-70-7
CN
DR
     Unspecified
MF
CI
     COM, MAN
                  ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO,
LC
     STN Files:
       CA, CABA, CANCERLIT, CAPLUS, CHEMCATS, CHEMLIST, CIN, CSCHEM, DDFU,
       DIOGENES, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IMSDRUGNEWS,
       IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK*, PROMT, RTECS*, TOXCENTER,
       USPATFULL, VETU
         (*File contains numerically searchable property data)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
             155 REFERENCES IN FILE CA (1907 TO DATE)
               4 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
             155 REFERENCES IN FILE CAPLUS (1907 TO DATE)
L16 ANSWER 5 OF 6 REGISTRY COPYRIGHT 2003 ACS on STN
RN (9041-08-1) REGISTRY
     Heparin, sodium salt (8CI, 9CI) (CA INDEX NAME)
OTHER NAMES:
     Alfa 87-120
CN
     Alfa 87-163
     Alfa 87-198
CN
     Alfa 87,-81
CN
CN
     Alfa 88-247
    (Ardeparin_sodium_)
CN
CN
     Bemiparin sodium
CN
     Clexan
CN
    (Dalteparin sodium )
CN
     Deligoparin sodium
CN
     Depo-Heparin
   Enoxaparin sodium
CN
CN
     Fragmin
CN
     Fragmin IV
CN
     H 2149
CN
    ·Hed-Heparin
CN
     Hepalean .
     Heparin Fragment Kabi 2165
CN
CN
     Heparin sodium
CN
     Hepathrom
CN
     Heprinar .
CN
     Hepsal
CN
     Inno-Hep
CN
     Kabi 2165
CN
     LHN 1
     Lioton 1000
CN
CN
     Lipo-Hepin
CN
     Lipo-Hepinette
CN
     Liquaemin sodium
     Liquemin
CN
CN
     Logiparin
CN
     Longheparin
CN
     Lovenox
CN
     Minihep
CN
     Minolteparin sodium
CN
     Monoparin
CN
     Normiflo
```

```
OP 2000
CN
CN
     Panheprin
CN
    Parnaparin-sodium-
CN
     PK 10169
CN
     Pularin
   Reviparin sodium
CN
CN
CN
     RP 54563
CN
     Sodium acid heparin
CN
     Sodium heparin
     Sodium heparinate
CN ·
     Sodium parnaparin
CN
     Thrombo-Hepin
CN
    (Tinzaparin sodium
CN
ADDITIONAL NAMES NOT AVAILABLE IN THIS FORMAT - Use FCN, FIDE, or ALL for
     DISPLAY
     12656-11-0, 101921-26-0, 102785-31-9
MF
     Unspecified
CI
     PMS, COM, MAN
PCT Manual registration, Polyester, Polyester formed
                ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS,
    STN Files:
LC
       BIOSIS, BIOTECHNO, CA, CABA, CAPLUS, CBNB, CHEMCATS, CHEMLIST, CIN,
       CSCHEM, DDFU, DETHERM*, DIOGENES, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB,
       IMSCOSEARCH, IMSDRUGNEWS, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK*,
       MSDS-OHS, NIOSHTIC, PHAR, PROMT, RTECS*, TOXCENTER, USAN, USPAT2,
       USPATFULL
         (*File contains numerically searchable property data)
     Other Sources: DSL**, TSCA**, WHO
         (**Enter CHEMLIST File for up-to-date regulatory information)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
            1111 REFERENCES IN FILE CA (1907 TO DATE)
              86 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            1114 REFERENCES IN FILE CAPLUS (1907 TO DATE)
L16 ANSWER 6 OF 6 REGISTRY COPYRIGHT 2003 ACS on STN
RN (9005=49=6. REGISTRY
    Heparin (8CI, 9CI) (CA INDEX NAME)
OTHER NAMES:
    .alpha.-Heparin
CN
CN
     Arteven
CN
     Bemiparin
   (Certoparin )
CN
CN
    Clevarin
    Clexane
CN
    Clivarin
CN
CN
    Clivarine
CN
    CY 216
CN
    CY 222
CN
   (Dalteparin)
CN
    (Enoxaparin )
CN
     Fluxum
CN
     FR 860
CN
    Fragmin A
CN
     Fragmin B
CN
     Fraxiparin
CN
     Heparin subcutan
CN
     Heparin sulfate
CN
    Heparinic acid
CN
    KB 101
CN
    Leparan
CN
    Mono-embolex
CN
    Multiparin
```

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Page 6

```
CN Nadroparin
Novoheparin
CN
     OP 386
CN
     OP 622
   · Pabyrn
     Parnaparin-
    Parvoparin
CN
    Reviparin-Sandoparin
CN
CN
CN
     Sublingula
CN
   CN
CN
     Vitrum AB
     9075-96-1, 11078-24-3, 11129-39-8, 104521-37-1, 37324-73-5, 91449-79-5
DR
MF
     Unspecified
     PMS, COM, MAN
CI
     Manual registration, Polyester, Polyester formed
PCT
     STN Files: ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO,
LC
       CA, CABA, CANCERLIT, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMLIST,
       CIN, CSCHEM, DDFU, DIOGENES, DRUGU, EMBASE, HSDB*, IFICDB, IFIPAT,
       IFIUDB, IMSCOSEARCH, IMSDRUGNEWS, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE,
       MRCK*, MSDS-OHS, NAPRALERT, NIOSHTIC, PHAR, PIRA, PROMT, RTECS*,
       TOXCENTER, USAN, USPAT2, USPATFULL
         (*File contains numerically searchable property data)
                      DSL**, EINECS**, WHO
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
           22491 REFERENCES IN FILE CA (1907 TO DATE)
            1806 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
           22518 REFERENCES IN FILE CAPLUS (1907 TO DATE)
```

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FILE COVERS 1907 - 12 Dec 2003 VOL 139 ISS 25 FILE LAST UPDATED: 11 Dec 2003 (20031211/ED)

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		•
L4	2	SEA FILE=REGISTRY ABB=ON ENOXAPARIN?/CN
L5.	2	SEA FILE=REGISTRY ABB=ON NADROPARIN?/CN
L6	2	SEA FILE=REGISTRY ABB=ON PARNAPARIN?/CN
L7	2	SEA FILE=REGISTRY ABB=ON REVIPARIN?/CN
L8	2	SEA FILE=REGISTRY ABB=ON DALTEPARIN?/CN
L9	2	SEA FILE=REGISTRY ABB=ON PARNAPARIN?/CN SEA FILE=REGISTRY ABB=ON PARNAPARIN?/CN SEA FILE=REGISTRY ABB=ON DALTEPARIN?/CN SEA FILE=REGISTRY ABB=ON TINZAPARIN?/CN SEA FILE=REGISTRY ABB=ON DANAPAROID/CN
L10	1	SEA FILE=REGISTRY ABB=ON DANAPAROID/CN
L11		SEA FILE-REGISTRY ADB-ON ARDEPARIN?/CN
L12	· . 1	SEA FILE=REGISTRY ABB=ON CERTOPARIN/CN
L13	1	SEA FILE=REGISTRY ABB=ON "CY 222"/CN
L14	ī	SEA FILE=REGISTRY ABB=ON "SR 90107"/CN
L15	ī	SEA FILE=REGISTRY ABB=ON "SR 90107"/CN SEA FILE=REGISTRY ABB=ON "ORG 31540"/CN
L16	. 6	SEA FILE=REGISTRY ABB=ON (L4 OR L5 OR L6 OR L7 OR L8 OR L9 OR
		L10 OR L11 OR L12 OR L13 OR L14 OR L15)
L17	23396	SEA FILE=CAPLUS ABB=ON L16
L18		SEA FILE=REGISTRY ABB=ON HEPARIN/CN
L19		SEA FILE=CAPLUS ABB=ON L18/D(L)LOW
. L20		SEA FILE=CAPLUS ABB=ON LATERAL SCLEROSIS
	892	SEA FILE=CAPLUS ABB=ON (SPINAL OR PROGRESSIVE OR INFANTILE) (1W
)MUSCULAR ATROPH?
L22	2	SEA FILE=CAPLUS ABB=ON (BULBO SPINAL OR BULBOSPINAL) (W) (NEUROP
L23	. 0	SEA FILE=CAPLUS ABB=ON MYELOPATHIC MUSCULAR ATROPH?
L24	26	SEA FILE=CAPLUS ABB=ON KUGELBERG (A) WELANDER OR WERDNIG (A) HOFFM
		AN
L25	690	SEA FILE=CAPLUS ABB=ON MOTOR(W) (SYSTEM OR NEURON) (2A) DISEASE#
L26	. 30	SEA FILE=CAPLUS ABB=ON BULBAR(W) (PALSY OR PARALYSIS)
L34	977	SEA FILE=CAPLUS ABB=ON ENOXAPARIN# OR NADROPARIN# OR PARNAPARI
	•	N# OR REVIPARIN# OR DALTEPARIN# OR TINZAPARIN# OR DANAPAROID#
		OR ARDEPARIN# OR CERTOPARIN# OR CY 222 OR SR 90107 OR ORG
		31540
L35	14-	SEA-FILE=CAPLUS ABB=ON (L17 OR L19 OR L34)-AND-(L20 OR L21 OR
		L22 OR L23 OR L24 OR L25 OR L26)
	-	The same of the sa

^{=&}gt; fil medl; d que 13

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```
FILE MEDLINE ENTERED AT 15:15:01 ON 12 DEC 2003
```

FILE LAST UPDATED: 2 DEC 2003 (20031202/UP). FILE COVERS 1958 TO DATE.

On April 13, 2003, MEDLINE was reloaded. See HELP RLOAD for details.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2003 vocabulary. See http://www.nlm.nih.gov/mesh/changes2003.html for a description on changes.

This file contains CAS Registry Numbers for easy and accurate substance identification.

```
L1 4315 SEA FILE=MEDLINE ABB=ON HEPARIN, LOW-MOLECULAR-WEIGHT+NT/CT

L2 15424 SEA FILE=MEDLINE ABB=ON MOTOR NEURON DISEASE+NT/CT - this term includes

13 0 SEA FILE=MEDLINE ABB=ON—L1—AND—L2—

The specific diseases

of claim 3
```

=> fil uspatf; d que 159

FILE 'USPATFULL' ENTERED AT 15:15:02 ON 12 DEC 2003
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FILE COVERS 1971 TO PATENT PUBLICATION DATE: 11 Dec 2003 (20031211/PD)
FILE LAST UPDATED: 11 Dec 2003 (20031211/ED)
HIGHEST GRANTED PATENT NUMBER: US6662368
HIGHEST APPLICATION PUBLICATION NUMBER: US2003229929
CA INDEXING IS CURRENT THROUGH 11 Dec 2003 (20031211/UPCA)
ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 11 Dec 2003 (20031211/PD)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Oct 2003
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Oct 2003

```
>>> USPAT2 is now available. USPATFULL contains full text of the
                                                                      <<<
                                                                       <.<<
>>> original, i.e., the earliest published granted patents or
>>> applications. USPAT2 contains full text of the latest US
                                                                       <<<
>>> publications, starting in 2001, for the inventions covered in
                                                                       <<<
>>> USPATFULL. A USPATFULL record contains not only the original
                                                                       <<<
                                                                       <<<
>>> published document but also a list of any subsequent
>>> publications. The publication number, patent kind code, and
                                                                       <<<
                                                                       <<<
>>> publication date for all the US publications for an invention
    are displayed in the PI (Patent Information) field of USPATFULL
                                                                       <<<
     records and may be searched in standard search fields, e.g., /PN,
                                                                       <<<
>>>
                                                                       <<<
>>> /PK, etc.
                                                                       <<<
     USPATFULL and USPAT2 can be accessed and searched together
>>>
     through the new cluster USPATALL. Type FILE USPATALL to
                                                                       <<< .
>>>
                                                                       <<<
     enter this cluster.
>>>
                                                                       ///
```

>>> Use USPATALL when searching terms such as patent assignees, <<< >>> classifications, or claims, that may potentially change from >>> the earliest to the latest publication. <<<

This file contains CAS Registry Numbers for easy and accurate substance identification.

L4	2 SE	A FILE=REGISTRY	ABB=ON	ENOXAPARIN?/CN
L5		A FILE=REGISTRY		NADROPARIN?/CN
				PARNAPARIN?/CN
L6		A FILE=REGISTRY		REVIPARIN?/CN
L7		A FILE=REGISTRY		
L8	2 SE	A FILE=REGISTRY	ABB=ON	DALTEPARIN?/CN

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```
L9
               2 SEA FILE=REGISTRY ABB=ON
                                            TINZAPARIN?/CN
L10
              1 SEA FILE=REGISTRY ABB=ON
                                            DANAPAROID/CN
L11
              1 SEA FILE=REGISTRY ABB=ON
                                            ARDEPARIN?/CN
L12
              1 SEA FILE=REGISTRY ABB=ON
                                            CERTOPARIN/CN
L13
                                            "CY 222"/CN
              1 SEA FILE=REGISTRY ABB=ON
L14
              1 SEA FILE=REGISTRY ABB=ON
                                            "SR 90107"/CN
L15
              1 SEA FILE=REGISTRY ABB=ON
                                            "ORG 31540"/CN
L16
               6 SEA FILE=REGISTRY ABB=ON
                                            (L4 OR L5 OR L6 OR L7 OR L8 OR L9 OR
                 L10 OR L11 OR L12 OR L13 OR L14 OR L15).
L49
           2233 SEA FILE=USPATFULL ABB=ON L16 OR (ENOXAPARIN# OR NADROPARIN#
                 OR PARNAPARIN# OR REVIPARIN#)/IT,TI,AB,CLM
L50
             26 SEA FILE=USPATFULL ABB=ON
                                             (DALTEPARIN# OR TINZAPARIN# OR
                 DANAPAROID# OR ARDEPARIN# OR CERTOPARIN#)/IT, TI, AB, CLM
L51
              5 SEA FILE=USPATFULL ABB=ON
                                             (CY 222 OR SR 90107 OR ORG 31540 OR
                 CALCIPARIN# OR HEPACARIN#)/IT, TI, AB, CLM
L52
            253 SEA FILE-USPATFULL ABB-ON
                                             (HEPARIN(3A)LOW)/IT,TI,AB,CLM
L53
                                             (LATERAL SCLEROSIS)/IT, TI, AB, CLM
           1269 SEA FILE=USPATFULL ABB=ON
L54
            133 SEA FILE=USPATFULL ABB=ON
                                             ((SPINAL OR PROGRESSIVE OR INFANTILE
                 OR MYELOPATHIC) (1W) MUSCULAR ATROPH?) / IT, TI, AB, CLM
L55
              O SEA FILE=USPATFULL ABB=ON
                                             ((BULBO SPINAL OR BULBOSPINAL) (W) (NE
                UROPATH? OR ATROPH?))/IT, TI, AB, CLM
L56
              6 SEA FILE-USPATFULL ABB-ON (KUGELBERG (A) WELANDER OR WERDNIG (A) H
                OFFMAN)/IT, TI, AB, CLM
L57
            104 SEA FILE=USPATFULL ABB=ON
                                             (MOTOR(W) (SYSTEM OR NEURON) (2A) DISEA
                SE#)/IT, TI, AB, CLM
L58
             8 SEA FILE=USPATFULL ABB=ON
                                             (BULBAR(W) (PALSY OR PARALYSIS))/IT, T
                1, AB, CLM
             13 SEA FILE=USPATFULL ABB=ON (L49 OR L50 OR L51-OR-L52)-AND (L53
159
                OR L54 OR L55 OR L56 OR L57 OR L58)
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=> fil embase; d que 164

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```
L4
             2 SEA FILE=REGISTRY ABB=ON
                                            ENOXAPARIN?/CN
 L5
               2 SEA FILE=REGISTRY ABB=ON
                                            NADROPARIN?/CN
 L6
               2 SEA FILE=REGISTRY ABB=ON
                                            PARNAPARIN?/CN
 1.7
               2 SEA FILE=REGISTRY ABB=ON
                                            REVIPARIN?/CN
 L8
              2 SEA FILE=REGISTRY ABB=ON
                                            DALTEPARIN?/CN
 L9
               2 SEA FILE=REGISTRY ABB=ON
                                            TINZAPARIN?/CN
 L10
               1 SEA FILE=REGISTRY ABB=ON
                                            DANAPAROID/CN
 L11
               1 SEA FILE=REGISTRY ABB=ON
                                            ARDEPARIN?/CN
 L12
               1 SEA FILE=REGISTRY ABB=ON
                                            CERTOPARIN/CN
 L13
               1 SEA FILE=REGISTRY ABB=ON
                                            "CY 222"/CN
 L14
               1 SEA FILE=REGISTRY ABB=ON
                                            "SR 90107"/CN
 L15
                                            "ORG -31540"/CN
               1 SEA FILE=REGISTRY ABB=ON
 L16
               6 SEA FILE=REGISTRY ABB=ON
                                            (L4 OR L5 OR L6 OR L7 OR L8 OR L9 OR .
                 L10 OR L11 OR L12 OR L13 OR L14 OR L15)
 L60
            5445 SEA FILE=EMBASE ABB=ON L16
 L61
           10267 SEA FILE=EMBASE ABB=ON LOW MOLECULAR WEIGHT HEPARIN+NT/CT
 L62
            9640 SEA FILE=EMBASE ABB=ON SPINAL MUSCULAR ATROPHY+NT/CT
 L63
             286 SEA FILE=EMBASE ABB=ON WERDNIG HOFFMANN DISEASE/CT
164خ
               5 SEA FILE=EMBASE ABB=ON_(L60_OR_L61) AND (L62 OR L63)
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=> fil PASCAL, JICST-EPLUS, ESBIOBASE, BIOTECHDS, LIFESCI, CONFSCI, WPIDS; d que 175

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L65 1190 SEA ENOXAPARIN# OR NADROPARIN# OR PARNAPARIN# OR REVIPARIN# L66 799 SEA ABB=ON DALTEPARIN# OR TINZAPARIN# OR DANAPAROID# OR ARDEPARIN# OR CERTOPARIN# L67 152 SEA ABB=ON CY 222 OR SR 90107 OR ORG 31540 OR CALCIPARIN#	3.
ARDEPARIN# OR CERTOPARIN#	R
152 SEA ARR=ON CY 222 OR SR 90107 OR ORG 31540 OR CALCIPARIN# 0	R
TO! TIS DEU UDD-ON OF SEE ON DO SEE ON ON ONE OF STEEL	
HEPACARIN#	
L68 5189 SEA ABB=ON HEPARIN(3A) LOW	
L69 11801 SEA ABB=ON LATERAL SCLEROSIS	
170 2215 SEA ABB=ON (SPINAL OR PROGRESSIVE OR INFANTILE OR MYELOPATH	IC)
(1W) MUSCULAR ATROPH?	
L71 9 SEA ABB=ON (BULBO SPINAL OR BULBOSPINAL) (W) (NEUROPATH? OR	
ATROPH?)	
L72 260 SEA ABB=ON KUGELBERG(A) WELANDER OR WERDNIG(A) HOFFMAN	
L73 3386 SEA ABB=ON MOTOR(W)(SYSTEM OR NEURON)(2A) DISEASE#	,
L74 367 SEA ABB=ON BULBAR(W) (PALSY OR PARALYSIS)	
(L75 1 SEA (L65 OR L66 OR L67 OR L68) AND (L69 OR L70 OR L71 OR L72	\supset
OR L73 OR L74)	•

=> fil DRUGU, BIOTECHNO, CABA, IPA, BIOSIS, TOXCENTER; d que 148

FILE 'DRUGU') ENTERED AT 15:15:04 ON 12 DEC 2003 COPYRIGHT (C) 2003 THOMSON DERWENT

(FILE_BIOTECHNO) ENTERED AT 15:15:04 ON 12 DEC 2003
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FILE 'IPA' ENTERED AT 15:15:04 ON 12 DEC 2003

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FILE BIOSIS; ENTERED AT 15:15:04 ON 12 DEC 2003

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Page 11

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(FILE 'TOXCENTER' ENTERED AT 15:15:04 ON 12 DEC 2003 COPYRIGHT (C) 2003 ACS

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2 SEA FILE=REGISTRY ABB=ON ENOXAPARIN?/CN
             2 SEA FILE=REGISTRY ABB=ON NADROPARIN?/CN
L5 ·
             2 SEA FILE=REGISTRY ABB=ON PARNAPARIN?/CN
L6
             2 SEA FILE=REGISTRY ABB=ON REVIPARIN?/CN
L7.
             2 SEA FILE=REGISTRY ABB=ON DALTEPARIN?/CN
rs
            2 SEA FILE=REGISTRY ABB=ON TINZAPARIN?/CN
L9
             1 SEA FILE=REGISTRY ABB=ON DANAPAROID/CN
L10
             1 SEA FILE=REGISTRY ABB=ON ARDEPARIN?/CN
L11
           1 SEA FILE=REGISTRY ABB=ON CERTOPARIN/CN
L12
             1 SEA FILE-REGISTRY ABB-ON "CY 222"/CN
L13
             1 SEA FILE=REGISTRY ABB=ON "SR 90107"/CN
L14
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                                          "ORG 31540"/CN
L15
              6 SEA FILE=REGISTRY ABB=ON (L4 OR L5 OR L6 OR L7 OR L8 OR L9 OR
L16
                L10 OR L11 OR L12 OR L13 OR L14 OR L15)
          61653 SEA L16 OR ENOXAPARIN# OR NADROPARIN# OR PARNAPARIN# OR
L38
                REVIPARIN#
           3469 SEA DALTEPARIN# OR TINZAPARIN# OR DANAPAROID# OR ARDEPARIN# OR
L39
                CERTOPARIN#
            742 SEA CY 222 OR SR 90107 OR ORG 31540 OR CALCIPARIN# OR HEPACARIN
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         14458 SEA HEPARIN(3A) LOW
L41
         11428 SEA LATERAL SCLEROSIS
L42
          3072 SEA ABB=ON (SPINAL OR PROGRESSIVE OR INFANTILE OR MYELOPATHIC)
L43
                (1W) MUSCULAR ATROPH?
              9 SEA ABB=ON (BULBO SPINAL OR BULBOSPINAL) (W) (NEUROPATH? OR
L44.
                ATROPH?)
            260 SEA ABB=ON KUGELBERG(A) WELANDER OR WERDNIG(A) HOFFMAN
           4122 SEA ABB=ON MOTOR(W) (SYSTEM OR NEURON) (2A) DISEASE#
L46
            384 SEA ABB=ON BULBAR (W) (PALSY-OR-PARALYSIS)-
L47
            3_SEA_(L38_OR_L39_OR-L40-OR-L41) AND (L42-OR_L43-OR-L44_OR_L45)
            OR L46 OR L47)
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=> dup rem 135,159,148,164,175

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FILE 'WPIDS' ENTERED AT 15:15:06 ON 12 DEC 2003 COPYRIGHT (C) 2003 THOMSON DERWENT PROCESSING COMPLETED FOR L35 PROCESSING COMPLETED FOR L59 PROCESSING COMPLETED FOR L48

10/644109

Page 12

PROCESSING COMPLETED FOR L64 PROCESSING COMPLETED FOR L75

L76 35 DUP REM L35 L59 L48 L64 L75 (1 DUPLICATE REMOVED) >
ANSWERS '1-14' FROM FILE CAPLUS

ANSWERS '1-14' FROM FILE CAPLUS
ANSWERS '15-27' FROM FILE USPATFULL
ANSWERS '28-29' FROM FILE BIOSIS
ANSWERS '30-34' FROM FILE EMBASE
ANSWER '35' FROM FILE WPIDS

=> d ibib ab hitrn 1-27; d iall 28-35;) fil hom

L76 ANSWER 1 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN DUPLICATE 1

ACCESSION NUMBER:

· 2002:283439 CAPLUS

DOCUMENT NUMBER:

137:936

TITLE:

Co-administration of IGF-I and glycosaminoglycans

greatly delays motor neuron

disease and affects IGF-I expression in the

wobbler mouse: A long-term study

AUTHOR(S):

Gorio, Alfredo; Lesma, Elena; Madaschi, Laura; Di

Giulio, Anna Maria

CORPORATE SOURCE:

Pharmacological Laboratories, Departments of Medicine, Surgery and Odontoiatry, Polo H San Paolo, Faculty of

Medicine, Milan, 20142, Italy

SOURCE:

Journal of Neurochemistry (2002), 81(1), 194-202

CODEN: JONRA9; ISSN: 0022-3042

PUBLISHER:

Blackwell Publishing Ltd.

DOCUMENT TYPE: LANGUAGE:

Journal English

The study on wobbler mouse has shown that the combined treatment with low doses of glycosaminoglycans (GAGs) and IGF-I fully prevented motor neuron death and forelimb impairment up to 9-12 wk of a mouse's life. The effect was accompanied by the prevention of the early hypertrophy of wobbler neurons, an effect likely due to the promotion of neuronal survival. At the 18th week, wobbler mice treated with IGF-I + GAGs still showed significantly improved forelimb function, reduced muscle atrophy and a higher no. of cervical motor neurons. IGF-I alone and GAGs alone were active up to the 3rd week of treatment; thereafter the beneficial effects of single treatments decreased drastically. GAGs and IGF-I treatments also affected IGF-I plasma and muscle levels. In wobbler mice there was a progressive redn. in IGF-I plasma levels that was prevented by IGF-I or GAGs alone and greatly increased, even above heterozygote levels, by the combination treatment. Such a powerful increase was correlated by a small enhancement in IGFBP-3 plasma levels, while treatment with IGF-I alone affected very significantly both IGFBP-1 and IGFBP-3. Co-treatment also prevented the decrease in IGF-I content obsd. in vehicle-treated wobbler mice forelimb muscles.

IT 9005-49-6, Heparin sulfate, biological studies

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(combined with dermatan sulfate; IGF-I and glycosaminoglycan

coadministration delays motor neuron

disease and affects IGF-I expression in wobbler mouse)
CE COUNT: 55 THERE ARE 55 CITED REFERENCES A

REFERENCE COUNT:

THERE ARE 55 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L76 ANSWER 2 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER:

2003:397015 CAPLUS

DOCUMENT NUMBER:

138:397896

TITLE:

Secreted Frizzled-related protein 1 inducing differentiation of embryonic stem cells into

ectodermal cells and its use

INVENTOR(S):

Sasai, Yoshiki; Iwata, Hiroo; Murakami, Yoshinobu;

Satoh, Mitsuo; Kobori, Masato; Yano, Keiichi

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Page 13

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PATENT ASSIGNEE(S):
```

Kyowa Hakko Kogyo Co., Ltd., Japan

SOURCE:

PCT Int. Appl., 156 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

```
APPLICATION NO.
PATENT NO.
                    KIND DATE
                                                                  DATE
                      A1
                            20030522
                                              WO 2002-JP11894 20021114
WO 2003042384
    W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
         CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
         GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
         LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
         PL, PT, RO, RU, SC, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT,
         TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ,
         MD, RU, TJ, TM
    RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
         NE, SN, TD, TG
                                           JP 2001-350724
                                                               A 20011115
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PRIORITY APPLN. INFO.:

A method of obtaining a soln. having an activity of inducing the differentiation of embryo stem cells into ectodermal cells or ectoderm-origin cells; and factors obtained by such method for inducing the differentiation of embryo stem cells into ectodermal cells or ectoderm-origin cells; are disclosed. The method involves the step of culturing stroma cells on a liq. medium contg. a polyanion compd. , neg. charged copolymer or homopolymer and then recovering the liq. medium. Also provided are: stroma cells or a factor derived from stroma cells possessing an activity to induce the differentiation in this method; cells induced by this method; and a method for increasing the purity of cells obtained by culturing in the presence of anticancer agents. A method is also provided for evaluating/screening a substance related to the regulation of the differentiation process from embryonic stem cells to ectodermal cells or cells derived from ectoderm by performing this method. Also provided are the pharmaceuticals contg. the above-described stroma cells or the factor derived from the stroma cells, or the above-described cells. The authors identified a stromal cell-derived inducing activity (SDIA), which induces differentiation of neural cells, including midbrain tyrosine hydroxylase-pos. (TH+) dopaminergic neurons, from mouse embryonic stem cells. The authors report here that SDIA is Secreted Frizzled-related protein 1 (SFRP1), a member of a protein family that contains a cysteine-rich domain similar to the WNT-binding site of Frizzled receptors and regulates the WNT pathway and induces efficient neural differentiation in embryonic stem cells.

9005-49-6, Heparin, biological studies

RL: BUU (Biological use, unclassified); MOA (Modifier or additive use); BIOL (Biological study); USES (Uses)

(culturing stroma cells on a liq. medium contg.; Secreted

Frizzled-related protein 1 inducing differentiation of embryonic stem cells into ectodermal cells and its use)

REFERENCE COUNT:

THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L76 ANSWER 3 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER:

2003:117964 CAPLUS

DOCUMENT NUMBER:

138:165523.

TITLE:

Hybrid proteins with neuregulin heparin-binding domain for targeting to heparan sulfate proteoglycans and

therapcutic uses thereof

INVENTOR(S):

Loeb, Jeffrey A.

10/644109

Page 14

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PATENT .ASSIGNEE(S):
```

Wayne State University, USA

SOURCE:

PCT Int. Appl., 74 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2003012045 A2 20030213 WO 2002-US24053 20020731

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.:

US 2001-308563P P 20010731

The present invention discloses that the neuregulin (NRG) heparin binding domain (N-HBD) functions to keep the EGF-like domain at sufficiently high concns. near erbB receptors for a sufficiently long period of time necessary to induce events downstream from receptor binding. In particular, fusion polypeptides are produced that comprise, as a targeting structure, a N-HBD polypeptide, fragment, homolog or functional deriv. and a protein to be targeted. This is fused to a polypeptide or peptide being targeted (Ptrg) to cell surfaces rich in heparan sulfate proteoglycans to either activate or inhibit interactions at tyrosine kinase receptors. Such products are used to treat diseases or conditions where either agonism or antagonism at tyrosine kinase receptors has beneficial effects, including cancer and a multitude of diseases of the nervous system. The present inventor examd. how NRG-HSPG interactions affect NRG-erbB receptor binding, erbB receptor auto- phosphorylation and downstream activation of AChR genes and newly-synthesized proteins in primary chick myotube cultures.

IT 9005-49-6, Heparin, biological studies

RL: BSU (Biological study, unclassified); BIOL (Biological study) (hybrid proteins with neuregulin heparin-binding domain for targeting to heparan sulfate proteoglycans and therapeutic uses thereof)

L76 ANSWER 4 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER:

2003:76905 CAPLUS

DOCUMENT NUMBER:

138:133498

TITLE:

Production of radial glial cells from neural stem cells and ependymal cells in the presence of growth

factors and therapeutic applications Weiss, Samuel; Gregg, Christopher

INVENTOR(S):
PATENT ASSIGNEE(S):

Stem Cell Therapeutics Inc., Can. PCT Int. Appl., 51 pp.

SOURCE: PCT Int. Appl

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2003008566 A1 20030130 WO 2002-CA1087 20020719

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,

Page 15

Krishnan 10/644109

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CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
             UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
             TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
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             PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
             NE, SN, TD, TG
                            20030213
                                         · US 2002-196549
                                                             20020717
     US 2003032181
                       A1
                                        US 2001-307096P P 20010720
CA 2001-2364095 A 20011130
PRIORITY APPLN. INFO .:
     The present invention relates to a method of producing radial glial cells
AB
     from neural stem cells, particularly by contacting neural stem cells with
     epidermal growth factor (EGF), fibroblast growth factor 2 (FGF-2) and/or
     TGF.alpha.. Leukemia inhibitory factor (LIF) and ciliary neurotrophic
     factor (CNTF) can optionally be added to enhance the effect of EGF, FGF-2
     or TGF.alpha.. Also provided are methods of producing radial glial cells
     from ependymal cells, as well as methods of proliferating ependymal cells.
     A method for treating a CNS disease in a mammal by transplanting radial
     glial cells into the mammal is disclosed. A method for enhancing neural
     cell mobilization in a mammal by administering a radial glia promoting
     agent ot transplanting radial glial cells into the mammal is also
     provided.
     9005-49-6, Heparin sulfate, biological studies
ľΤ
     RL: BSU (Biological study, unclassified); BUU (Biological use,
     unclassified); BIOL (Biological study); USES (Uses)
        (glial cells prodn. in presence of growth factors and; prodn. of radial
        glial cells from neural stem cells and ependymal cells in presence of
        growth factors and therapeutic applications)
                               THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS
                         7
REFERENCE COUNT:
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L76 ANSWER 5 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN
ACCESSION NUMBER:
                         2003:6095 CAPLUS
                         138:52352
DOCUMENT NUMBER:
                         Method of producing region-specific neurons from human
TITLE:
                         neuronal stem dells
                         Wu, Ping F.
INVENTOR(S):
                         The University of Texas System, USA
PATENT ASSIGNEE(S):
                         PCT Int. Appl., 62 pp.
SOURCE:
                         CODEN: PIXXD2
                         Patent
DOCUMENT TYPE:
                         English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
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KIND DATE
                                          APPLICATION NO.
    PATENT NO.
                                          WO 2002-US19743 20020619
                      A2
                            20030103
    WO 2003000852
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, M2, NO, NZ, OM, PH,
            PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
            UA, UG, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,
            CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,
            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                           US 2002-176971 20020619
    US 2003013193
                      A1
                            20030116
                                        US 2001-300344P P 20010622
PRIORITY APPLN. INFO.:
    The invention concerns a method of priming neural stem cells in vitro by
```

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Page 16

adhesively culturing in a mixt. of basic fibroblast growth factor, laminin and heparin to differentiate into specific neuronal phenotypes, including cholinergic, glutamatergic and GABAergic neurons, in a region-specific manner, when transplanted in vivo.

IT 9005-49-6, Heparin, biological studies

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(method of producing region-specific neurons from human neuronal stem cells)

.L76 ANSWER 6 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER:

· 2003:588647 CAPLUS

DOCUMENT NUMBER:

139:112759

TITLE:

Protein and cDNA sequences of a human glial-derived

neurotrophic factor and therapeutic use for nervous

system diseases

INVENTOR(S):

Zhou, Shoushan; Zheng, Zanshun; Fang, Haizhou; Chen, Yong; Jiang, Ruofeng; Zhu, Aitang; Zhang, Qi; Gan,

Shuyan; Lan, Xuan

PATENT ASSIGNEE(S):

Zhuhai Yisheng Biopharmaceuticals Co., Ltd., Peop.

Rep. China

SOURCE:

Faming Zhuanli Shenqing Gongkai Shuomingshu, 28 pp.

CODEN: CNXXEV

DOCUMENT TYPE: LANGUAGE: Patent Chinese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE		APPLICATION NO.	DATE
CN 1364812	A	20020821	•	CN 2001-107450	20010111
PRIORITY APPLA, INFO	. •	•	C	N 2001-107450	20010111

AB The invention provides the DNA sequence or its DNA fragments and their encoded amino acid sequences of human glial-derived neurotrophic factor (GDNF) cloned from human glioma cell line C6 or synthesized by solid-phase synthesis method. The invention relates to the construction of the expression vector, the expression of GDNF in E.coli, yeast, and CHO cells, and sepn. and purifn. of GDNF from the cultured products of the above genetically engineered bacteria or CHO cells. The invention also relates to the application of the expressed GDNF in prepg. the medical compn. (composed of GDNF, natural ganglioside or its deriv., and/or mycose or hyaluronic acid) for treating nervous system disease, insanity, etc.

IT 9005-49-6, Heparin, uses

RL: DEV (Device component use); USES (Uses)

(protein and cDNA sequences of human glial-derived neurotrophic factor and therapeutic use for nervous system diseases)

L76 ANSWER 7 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER:

2001:851346 CAPLUS

DOCUMENT NUMBER:

135:368940

TITLE:

Novel method for inducing the differentiation of

embryonic stem cells into ectodermal cells and its use

INVENTOR(S): Sasai, Yoshiki; Nishikawa, Shinichi
PATENT ASSIGNEE(S): Kyowa Hakko Kogyo Co., Ltd., Japan

SOURCE:

PCT Int. Appl., 138 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

10/644109

Page 17

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WO 2001088100
                         A1
                              20011122
                                               WO 2001-JP4080
                                                                20010516
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
              CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
              GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
              LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT,
         RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
              BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                        A5
                              20011126
                                              AU 2001-56767
                                                                  20010516
     AU 2001056767
     US 2002151056
                              20021017
                                               US 2001-855587
                         A1
                                                                  20010516
                                               EP 2001-930185
     EP 1302533
                         A1
                              20030416
                                                                  20010516
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
              IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
                                                                 20000516
PRIORITY APPLN. INFO.:
                                            JP 2000-144059
                                            JP 2000-290819
                                                              Α
                                                                 20000925
                                           :US 2000-257049P P
                                                                  20001220
                                            WO 2001-JP4080 . W
                                                                 20010516
     A novel method for inducing the differentiation of embryonic stem cells
AB
     into ectodermal cells or cells derived from ectoderm is provided, in which
     a process for culturing embryonic stem cells in a non-aggregated state is
     included. Also provided are: culture medium and culture supernatant used
     for this method; a differentiation inducer used in this method; stroma
     cells or a factor derived from stroma cells possessing an activity to
     induce the differentiation in this method; an antibody capable of
     specifically recognizing the stroma cells; an antigen capable of
     recognizing the antibody; and cells induced by this method. A method is
     also provided for evaluating/screening a substance related to the
     regulation of the differentiation process from embryonic stem cells to
     ectodermal cells or cells derived from ectoderm by performing this method.
     Also provided are the pharmaceuticals contg. the above-described stroma
     cells or the factor derived from the stroma cells, the above-described
     antibody, the above-described antigen, or the above-described cells.
```

T . 9005-49-6, Heparin, biological studies

RL: BSU (Biological study, unclassified); BIOL (Biological study) (novel method for inducing differentiation of embryonic stem cells into ectodermal cells and use)

REFERENCE COUNT:

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L76 ANSWER 8 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER:

2000:824499 CAPLUS

DOCUMENT NUMBER:

134:14946

A-form of cytoplasmic domain of nARIA (CRD-neuregulin)

and uses in diagnosis and maintaining synaptic

connections

INVENTOR(S):

TITLE:

Role, Lorna W.; Talmage, David; Bao, Jianxin-

PATENT ASSIGNEE(S): The Trustees of Columbia University In the City of New

York, USA

SOURCE:

PCT Int. Appl., 210 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

			•	
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000070322	A2	20001123	WO 2000-US13157	20000512
WO 2000070322	A3	20011011		•
WO 2000070322	C2	20020926		
W. AF AT.	AM. AT	. AU. AZ. 1	RA. BR. RG. BR. BY. CA	CH. CN. CR.

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CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,
             IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA,
            MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,
                        TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ,
             SĶ, SL, TJ,
             BY, KG, KZ, MD, RU, TJ,
                                    TM
        RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
            DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
            CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
    AU 2000048474
                            20001205
                                           AU 2000-48474
                                                            20000512
                      A5
                                        US 1999-312596
PRIORITY APPLN: INFO .:
                                                        A 19990514
                                        WO 2000-US13157 W 20000512
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This invention provides an assay for diagnosing whether a subject has or is predisposed to developing a neoplastic disease which comprises: (a) obtaining a biol. sample from the subject; (b) contacting the sample with an agent that detects the presence of an extracellular domain of nARIA (CRD-neuregulin) or an isoform thereof; (c) measuring the amt. of agent bound by the sample; (d) comparing the amt. of agent bound measured in step (c) with the amt. of agent bound by a std. normal sample, a higher amt. bound by the sample from the subject being indicative of the subject having or being predisposed to developing a neoplastic disease. One embodiment of this invention is a method for maintaining synaptic connections between a neuron and a target cell comprising contacting the target cell with an nARIA polypeptide or a nucleic acid mol. encoding nARIA in an amt. sufficient to induce the formation of a synaptic junction.

IT 9005-49-6, Heparin sulfate, biological studies RL: BSU (Biological study, unclassified); BIOL (Biological study) (nARIA vs. ARIA affinity for; A-form of cytoplasmic domain of nARIA (CRD-neuregulin) and uses in diagnosis and maintaining synaptic connections)

L76 ANSWER 9 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 1998:744967 CAPLUS

DOCUMENT NUMBER: 130:839

Compositions and methods of therapy for TITLE:

IGF-I-responsive conditions

INVENTOR(S): Scharschmidt, Bruce F.; Gorio, Alfredo; Muller,

Eugenio E.

PATENT ASSIGNEE(S): Chiron Corp., USA

SOURCE: PCT Int. Appl., 36 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	ENT	NO.		KI	DI	DATE								DATE			,
WO	9850	062		A	1	1998	1112		W	0 19	98-U	5927	3	19980	0506		
	W:	AL,	AM,	AT,	ΛT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CU,	CZ,
		CZ,	DE,	DE,	DK,	DK,	EE,	EE,	ES,	FI,	FI,	GB,	GE,	GH,	GM,	GW,	HU,
		ID,	IL,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,
		MD,	MG,	MK,	MN,	MW,	MX,	NO,	NZ,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,
		SK,	SK,	SL,	TJ,	TM,	TR,	TT,	UA,	UG,	ŲS,	UZ,	VN,	ΥU,	ZW,	AM,	AZ,
		BY,	KG,	ΚZ,	MD,	RU,	TJ,	TM	•								
	RW:	GH,	GM,	KE,	LS,	MW,	SD,	SZ,	UG,	ZW,	AT,	BE,	CH,	CY,	DΕ,	DK,	ES,
		FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	CI,
		CM,	GA,	GN,	ML,	MR,	NÉ,	SN,	TD,	TG							
ΑU	9873										98-7	3707		1998	0506		
EΡ	1015	019		A	1	2000	0705		E	P 19	98-9	2100	4 ·	1998	0506	•	
														NL,			PT,
		IE,								•							
JΡ	2002	5071	93	T.	2	2002	0305		J.	P 19	98-5	4846	7	1998	0506		

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20021024
                                          US 2002-279343
    US 2003100505
                            20030529
                                       IT 1997-MI1042
                                                       A 19970506
PRIORITY APPLN. INFO.:
                                       WO 1998-US9273
                                                        W 19980506
                                       US 1999-423161
                                                        B1 19991101
    Compns. and methods useful in therapy for IGF-I (insulin-like growth
     factor-I)-responsive conditions in a mammal are provided. The method
     comprises concurrent thorapy with both IGF-I or a variant thereof and at
    least one GAG to promote a desired therapeutic response with respect to a
    particular IGF-I- responsive condition. Concurrent therapy is achieved by
     administering to a mammal a single pharmaceutical compn. contg. both IGF-I
     (or a variant thereof) and at least one GAG according to a dosing regimen.
    Alternatively, IGF-I or a variant thereof and at least one GAG can be
    administered as two sep. pharmaceutical compns. A pharmaceutical compn.
     comprising IGF-I or a variant thereof and at least one GAG for use in the
     IGF-I and GAG therapy is also provided. In expts. it was shown that
     compns. of rhIGF-I and glucosaminoglycans are effective in promoting
     desired therapeutic treatment effects in the animal model of ALS and
     spinal muscular atrophy.
     9005-49-6, Heparin, biological studies
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); MOA (Modifier or additive use); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (therapy for insulin like growth factor-I-responsive conditions)
                               THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                         5
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L76 ANSWER 10 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN
                         1998:485092 CAPLUS
ACCESSION NUMBER:
                         129:104688
DOCUMENT NUMBER:
                         Methods and substances for elevating the concentration
TITLE:
                         of free insulin-like growth factor in vivo, and
                         methods for screening the substances for clinical use
                         Sakano, Katsuichi; Higashihashi, Nobuyuki; Hashimoto,
INVENTOR(S):
                         Ryuji
                         Daiichi Pharmaceutical Co., Ltd., Japan
PATENT ASSIGNEE(S):
                         PCT Int. Appl., 67 pp.
SOURCE:
                         CODEN: PIXXD2
                         Patent
DOCUMENT TYPE:
                         Japanese
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                                          APPLICATION NO. DATE
                      KIND DATE
     PATENT NO.
                                         . _____
                            19980709
                                          WO 1997-JP4881
                                                           19971226
                  · A1
     WO 9829451
         W: AL, AU, BA, BB, BG, BR, CA, CN, CU, CZ, EE, GE, GM, GW, HU, ID,
             IL, IS, JP, KG, KR, LC, LK, LR, LT, LV, MG, MK, MN, MX, NO, NZ,
             PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, AM, AZ, BY,
             KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI,
             FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM,
             GA, GN, ML, MR, NE, SN, TD, TG
                                                            19971226
                            19980731
                                           AU 1998-78910
                      A1
     AU 9878910
                                                            19971226
                                          EP 1997-949250
                       A1
                            19991222
     EP 965596
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
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US 1999-331851 Disclosed are (1) methods of increasing the in vivo concn. of insulin-like AB

19990827

20020806

20030911

В1

A1

NO 9902785

US 6428781

US 2003170240

PRIORITY APPLN. INFO .:.

NO 1999-2785

JP 1996-349968

WO 1997-JP4881

US 1999-331851

US 2002-173805

19990608

19990628

20020619

A 19961227

W 19971226

A3 19990628

growth (IGF) by freeing the IGF from the IGF-IGFBP (IGF binding protein), (2) method of increasing the in vivo concn. of IGF-IGFBP concn. from the IGF-IGFBP-ALS (acid labile subunit) complex, and (3) methods for screening the substances that increase the in vivo concn. of IGF or IGF-IGFBP from their resp. precursors. Among 34 chem. compds. tested in vitro, ellagic acid, aclacinomycin A, and heparin were most effective on inhibiting the binding between IGF-II and IGFBP 3. Both human IGF-II[27-Tyr.fwdarw.Leu, 43-Val.fwdarw.Leu) and rabbit anti-rat IGFBP 3 were used to demonstrated their ability to increase the free IGF-I blood level in SD. rats. The substances are useful as a prophylactics or therapeutics for diseases, e.g., diabetes, amyotrophic lateral sclerosis , and osteoporosis, that can be treated by IGF.

9005-49-6, Heparin, biological studies

RL: ANT (Analyte); BSU (Biological study, unclassified); BUU (Biological use, unclassified); ANST (Analytical study); BIOL (Biological study); USES (Uses)

(IGF-II-IGFBP 3 binding inhibition by; methods and substances for elevating concn. of free insulin-like growth factor in vivo, and methods for screening substances for clin. use)

REFERENCE COUNT:

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L76 ANSWER 11 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN

8

ACCESSION NUMBER:

1998:197424 CAPLUS

DOCUMENT NUMBER:

128:266268

TITLE:

Identification of agents that protect against

inflammatory injury to neurons

INVENTOR(S):

Giulian, Dana J.

PATENT ASSIGNEE(S):

Baylor College of Medicine, USA; Giulian, Dana J.

SOURCE:

PCT Int. Appl., 149 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLICATION NO.	DATE
WO 9811923	A1 19980326	WO 1997-US16999	19970919
W: AU, CA,		TT	
US 6071493	A 20000606	FI, FR, GB, GR, IE; IT, US 1996-717551	
US 6043283	A 20000328	US 1997-870967	19970606
AU 9745894 AU 738509		AU 1997-45894	19970919
EP 1051195	A1 20001115	EP 1997-944385	
R: AT, BE, IE, FI	CH, DE, DK, ES,	FR, GB, GR, IT, LI, LU,	NL, SE, MC, PT,
JP 2002504988			
PRIORITY APPLN. INFO.	.:	US 1996-717551 A2 US 1997-870967 A2	

WO 1997-US16999 W 19970919 OTHER SOURCE(S): MARPAT 128:266268

Methods are disclosed for identifying agents that inhibit the toxic effects of neurotoxins on neurons from plaque component-activated mononuclear phagocytes. Also disclosed are methods for identifying agents that inhibit mononuclear phagocyte-plaque component complex formation, plaque component activation of mononuclear phagocytes, and plaque component-induced neurotoxicity of mononuclear phagocytes. The invention is also directed to agents and pharmaceutical compns. obtained by the identification methods described. Addnl., the invention describes methods for using tyramine compds. to inhibit the toxic effects of neurotoxins and methods to treat and diagnose neurodegenerative diseases and disorders.

Krishnan 10/6

TT 9005-49-6, Heparin sulfate, biological studies

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL

(Biological study); PROC (Process)

(identification of agents that protect against inflammatory injury to

neurons)

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L76 ANSWER 12 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER:

1998:180841 CAPLUS

DOCUMENT NUMBER:

128:239488

TITLE:

Polydithiocarbamate-containing macromolecules and the

use thereof for therapeutic and diagnostic

applications

INVENTOR(S):

Lai, Ching-San

PATENT. ASSIGNEE (S):

Medinox, Inc., USA; Lai, Ching-San

SOURCE:

PCT Int. Appl., 68 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PAT	ENT 1	NO.		ĶΙΙ	ND	DATE				APPL	ICAT	ION N	٥.	DATE	•		
	WO	98110				 1	1998	0319		i	WO 1	 997 <i>-</i> 1	us153	 24	1997	0828		
	110	W:	AT.	AM.	AT.	AIJ.	AZ.	BA.	BB.	BG	, BR	, BY	, CA,	CH.	CN,	CU,	CZ,	DE,
		***	DK.	EE.	ES.	FI.	GB.	GE.	GH.	HU	IL	. IS	, JP,	KE	KG,	KP,	KR,	KZ,
			LC.	I.K.	I.R.	LS.	LT.	LU.	LV.	MD	. MG	. MK	, MN,	MW	MX.	NO,	NZ,	PL,
			DT	BO.	RII.	SD.	SE.	SG.	SI.	SK	. SL	. TJ	, TM,	TR.	TT.	UA.	UG,	US,
													, MD,				•	•
		DW.	GH,	KF	LS	MW.	SD.	57.	IIG.	7.W	, AT	. BE	, СН,	DE.	DK.	ES.	FI.	FR.
		Ku.	GII,	CP,	TF.	ΤΨ.	LII.	MC.	NI.	PΨ	SE	, BF	, BJ,	CF.	CG.	ĊI.	CM.	GA.
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	7117	0741	725	LILI,	1317	1	1998	0402	10		AII 1	997-	41725		1997	0828		
		74679					2002			•	, i O I	,,,	11,20			0020		
	AU	0271	50 50		ע .	<u>.</u> 1	1002	0707			FD 1	997-	93969	Δ	1997	0828	•	
	EP	92/1:	3 M	חפ	CU.	DE.	エンフン	U/U/	ED.	CR	כם ביני	777 TT	, LI,	T.IT	NI.	SE	MC	PΤ.
		к:			Cn,	UE,	DR,	ES,	r rv,	GD	, 61	, 11	, 11,	ъод	,		1107	,
•			IE,	r I			1000				CN 1	007_	19779	7	1007	0828		
	CN	1230.	T 18	~ ~	A	_	1999	0323		•	UN I	000-	51368	۵.	1007	0020		
	JP	2002	2118	31	T.	Z .	2002	0410							1999			•,
		2000					2000	0626					7001 ₉					
PRIOF	RITY	(APP	LN.	INFO	.:								67P					
													087					
													67					
		•	•	•				•		WO	1997	-US1	5324	. W	1997	0828		

OTHER SOURCE(S): MARPAT 128:239488

A new class of drugs is provided for therapeutic treatment of such indications as cerebral stroke and other ischemia/reperfusion injury. Dithiocarbamates are linked to the surface of a macromol. (e.g. albumin), either by using crosslinking reagents or by non-specific binding, to produce polydithiocarbamate-macromol.-contg. compns. Combination therapeutic methods have been developed for the in vivo inactivation or inhibition of formation (either directly or indirectly) of species which induce the expression of inducible nitric oxide synthase, as well as reducing nitric oxide levels produced as a result of NO synthase expression. Magnetic resonance imaging methods have been developed for the measurement of cerebral and cardiac blood flow and infarct vol. in ischemic stroke or heart attack situations. Such methods employ iron-contq. complexes of a compn. comprising a dithiocarbamate and a macromol. as contrast agents. Prepn. of a reaction product of bovine serum albumin with N-methyl-D-glucamine dithiocarbamate is described. 9005-49-6D, Heparin, dithiocarbamate reaction products, biological

. . . .

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications)

REFERENCE COUNT:

THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L76 ANSWER 13 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER:

1996:394181 CAPLUS

DOCUMENT NUMBER:

125:49359

TITLE:

Use of receptor agonists to stimulate superoxide

dismutase activity

INVENTOR(S):

Marklund, Stefan L.; Straalin, Pontus

PATENT ASSIGNEE(S):

Swed.

SOURCE:

PCT Int. Appl., 70 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.			KIND DATE			APPLICATION NO.				ο.	DATE						
WO	WO 9614060		A1 19960517		WO 1995-IB979				19951103								
	W:	ΑM,	AT,	AT,	ΑU,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CZ,	CZ,	DE,	DE,	DK,
		DK,	EE,	EE,	ES,	FI,	FI,	GB,	GE,	HU,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,
		LK,	LR,	LT,	LU,	LV,	MD,	MG,	MK,	MN,	MW,	MX,	NO,	NZ,	PL,	PT,	RO,
		RU,															
	RW:	KE,	LS,	MW,	SD,	SZ,	υĜ,	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IE,
						PT,											
		NE,	SN,	TD,	TG												
AU	9537	082		Α	1	1996	0531		٠A١	J 19	95-3	7082		1995	1103		
PRIORIT	Y APP	LN.	INFO	.:					DK 1	994-	1283			1994	1104		
								1	WO 1	995-	IB97	9		1995:	1103		

The present invention relates to the use of a substance for the manuf. of AB a compn. for stimulating the release of extracellular superoxide dismutase (EC-SOD) from cells or stimulating the synthesis of EC-SOD in cells. In particular, the invention relates to the use of a substance for the manuf. of a compn. for prophylaxis or treatment of a disease or disorder connected with the presence of formation of superoxide radicals and other toxic intermediates derived from the superoxide radical. Further, the invention relates to a method for detq, the effect of a substance with respect to stimulating the release of EC-SOD from cells or stimulating the synthesis of EC-SOD in cells and to substances which have been selected by the method. Within the scope of the invention is a method of preventing, diminishing, controlling, or inhibiting a disease or disorder connected with the presence or formation of superoxide radicals and other toxic intermediates derived from the superoxide radical in a patient who has been established to have a high risk of developing a such disease or disorder, or who has developed such a disease or disorder, the method comprising administering an effective amt. of a substance which is capable of stimulating the release of EC-SOD from cells or stimulating the synthesis of EC-SOD in cells. SOD isoenzyme levels were detd. for a variety of human tissues and for the blood vessel wall of man and other mammals. Also reported is the reaction of cultured cells to a variety of factors (inflammation-related substances, vasoactive substances, growth factors, etc.).

IT 9005-49-6, Heparin

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

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Page 23

(superoxide dismutase stimulation with receptor agonists and therapeutic use)

L76 ANSWER 14 OF 35 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER:

1991:574647 CAPLUS

DOCUMENT NUMBER:

115:174647

TITLE:

Inhibition of cell growth by keratin sulfate, chondroitin sulfate, dermatan sulfate, and other

proteoglycans

INVENTOR(S):

Snow, Diane M.; Silver, Jerry; Harel, Adrian; Roufa,

PATENT ASSIGNEE(S):

Case Western Reserve University, USA; Gliatech, Inc.

SOURCE:

PCT Int. Appl., 133 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PATENT NO.	KIND DATE	APPLICATION NO.	DATE
	WO 9106303	A1 19910516	WO 1990-US6189	19901026
			ES, FI, HU, JP, KR, LK,	
		O, SE, SU		a. an an
			CH, CM, DE, DK, ES, FR,	GA, GB, GR, IT,
	LU, MI	L, MR, NL, SE, SN,	TD, TG	10001026
			CA 1990-2071898	
		A1 19910531	AU 1991-68726	19901026
	EP 493533		EP 1990-917627	
			FR, GB, IT, LI, LU, NL,	
	JP 06502840	T2 19940331		19901026 ·
F	PRIORITY APPLN. IN	FO.:	US 1989-428374	19891027
			WO 1990-US6189	19901026

Proteoglycans such as keratan sulfate (I), chondroitin sulfate (II), AΒ dermatan sulfate (III), heparan sulfate (IV), heparin (V), and hyaluronic acid (VI) are used to prevent neurite outgrowth, i.e. axonal growth, or nerve regeneration, or glial cell migration, invasion, or regeneration. Inhibitors and antagonists of proteoglycans may also be used to promote nerve growth or glial cell migration or invasion. Such inhibitors and antagonists include antibodies, degradative enzymes, lectins, and disaccharide antagonists of the receptors for I, II, III, IV, V, or VI. Chick E-6 dorsal root ganglia (DRG) cells were cultured on nitrocellulose treated with a II-proteoglycan in the presence of nerve growth factor. DRG neurite outgrowth was completely inhibited by 0.4 mg/mL II-proteoglycan.

9005-49-6D, Heparin, derivs. RL: BIOL (Biological study)

. (neurite outgrowth inhibition by)

L76 ANSWER 15 OF 35 USPATFULL on STN.

ACCESSION NUMBER:

2003:243831 USPATFULL

TITLE:

Composition of an endogenous insulin-like growth

factor-II derivative

INVENTOR(S):

Sakano, Katsuichi, Tokyo, JAPAN

Searched by Barb O'Bryen, STIC 308-4291

Higashihashi, Nobuyuki, Tokyo, JAPAN

PATENT ASSIGNEE(S):

Hashimoto, Ryuji, Tokyo, JAPAN DAIICHI PHARMACEUTICAL CO., LTD. (non-U.S. corporation)

·	NUMBER	KIND	DATE	
•				•
PATENT INFORMATION:	US 2003170240	A1	20030911	
APPLICATION INFO.:	US 2002-173805			
RELATED APPLN. INFO.:	Division of Ser.	No. US	1999-3318	51, filed on 28 Jun

10/644109

Page 24

1999, GRANTED, Pat. No. US 6428781 A 371 of

International Ser. No. WO 1997-JP4881, filed on 26 Dec

1997, UNKNOWN

NUMBER DATE

PRIORITY INFORMATION:

JP 1996-349968

19961227

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

SUGHRUE MION, PLLC, 2100 Pennsylvania Avenue, NW,

Washington, DC, 20037-3213

NUMBER OF CLAIMS:

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

12 Drawing Page(s)

LINE COUNT:

1522

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The effects of endogenous insulin-like growth factor can be appreciated by administering compounds capable of increasing free IGF in living bodies. Compounds are described which can elevate the concentration of unbound IGF by converting endogenous IGF (insulin-like growth factor) into free, biologically active IGF or elevating the concentration of the complex of IGF and IGFBP (insulin-like growth factor binding protein) in living bodies. Medicaments can be prepared containing these compounds or these compounds may be used in methods for the prevention and or treatment of IGF-responsive diseases such as diabetes, amyotrophic lateral sclerosis, or osteoporosis.

9005-49-6, Heparin, biological studies ΙT

(IGF-II-IGFBP 3 binding inhibition by; methods and substances for elevating concn. of free insulin-like growth factor in vivo, and methods for screening substances for clin. use)

L76 ANSWER 16 OF 35 USPATFULL on STN

ACCESSION NUMBER:

2003:146758 USPATFULL

TITLE:

Compositions and methods of therapy for

IGF-I-responsive conditions

INVENTOR(S):

Scharschmidt, Bruce F., San Francisco, CA, UNITED

STATES

Gorio, Alfredo, Milano, ITALY Muller, Eugenio E., Milano, ITALY

PATENT ASSIGNEE(S):

Chiron Corporation, Emeryville, CA, 94608 (U.S.

corporation)

NUMBER KIND PATENT INFORMATION: US 2003100505 A1 20030529 APPLICATION INFO.: US 2002-279343 Al 20021024 (10)

RELATED APPLN. INFO.: Continuation of Ser. No. US 1999-423161, filed on 1 Nov

1999, ABANDONED A 371 of International Ser. No. WO

1998-US9273, filed on 6 May 1998, PENDING

DOCUMENT TYPE: Utility FILE SEGMENT: LEGAL REPRESENTATIVE:

APPLICATION

Chiron Corporation, 4560 Horton Street, Emeryville, CA,

94608

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

13 1

NUMBER OF DRAWINGS:

1 Drawing Page(s)

LINE COUNT:

1002

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Compositions and methods useful in therapy for IGF-I-responsive conditions in a mammal are provided. The method comprises concurrent therapy with both IGF-I or a variant thereof and at least one GAG to promote a desired therapeutic response with respect to a particular IGF-I-responsive condition. Concurrent therapy is achieved by

Krishnan 10/644109 Page 25

administering to a mammal a single pharmaceutical composition containing both IGF-I (or a variant thereof) and at least one GAG according to a dosing regimen. Alternatively, IGF-I or a variant thereof and at least one GAG can be administered as two separate pharmaceutical compositions. A pharmaceutical composition comprising IGF-I or a variant thereof and at least one GAG for use in the IGF-I and GAG therapy is also provided.

IT 9005-49-6, Heparin, biological studies

(therapy for insulin like growth factor-I-responsive conditions)

L76 ANSWER 17 OF 35 USPATFULL on STN

ACCESSION NUMBER:

2003:44871 USPATFULL

TITLE: INVENTOR(S): Production of radial glial cells Weiss, Samuel, Calgary, CANADA

PATENT ASSIGNEE(S):

Gregg, Christopher, Calgary, CANADA Stem Cell Therapeutics Inc., Calgary, AB, CANADA

(non-U.S. corporation)

NUMBER DATE

PRIORITY INFORMATION:

CA 2001-2364095 · 20011130 US 2001-307096P 20010720 (60)

DOCUMENT TYPE: Util:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

BURNS DOANE SWECKER & MATHIS L L P, POST OFFICE BOX

1404, ALEXANDRIA, VA, 22313-1404

NUMBER OF CLAIMS: 2

EXEMPLARY CLAIM:

3 Drawing Page(s)

NUMBER OF DRAWINGS: LINE COUNT:

1123

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to a method of producing radial glial cells from neural stem cells, particularly by contacting neural stem cells with epidermal growth factor (EGF), fibroblast growth factor 2 (FGF-2) and/or TGF.alpha.. Leukemia inhibitory factor (LIF) and ciliary neurotrophic factor (CNTF) can optionally be added to enhance the effect of EGF, FGF-1 or TGF.alpha.. Also provided are methods of producing radial glial cells from ependymal cells, as well as methods of proliferating ependymal cells.

IT 9005-49-6, Heparin sulfate, biological studies

(glial cells prodn. in presence of growth factors and; prodn. of radial glial cells from neural stem cells and ependymal cells in presence of growth factors and therapeutic applications)

L76 ANSWER 18 OF 35 USPATFULL on STN

ACCESSION NUMBER:

2003:17441 USPATFULL

TITLE:

Method of producing region-specific neurons from human

neuronal stem cells

INVENTOR(S):

Wu, Ping, League City, TX, UNITED STATES

NUMBER DATE

PRIORITY INFORMATION:

US 2001-300344P 20010622 (60)

DOCUMENT TYPE:

Utility .

10/644109

Page 26

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

MUETING, RAASCH & GEBHARDT, P.A., P.O. BOX 581415,

MINNEAPOLIS, MN, 55458

NUMBER OF CLAIMS:

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

5 Drawing Page(s)

LINE COUNT:

1375

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method of priming neural stem cells in vitro by adhesively culturing in a mixture of basic fibroblast growth factor, laminin and heparin to differentiate into specific neuronal phenotypes, including cholinergic, glutamatergic and GABAergic neurons, in a region-specific manner, when

transplanted in vivo.

IT 9005-49-6, Heparin, biological studies

> (method of producing region-specific neurons from human neuronal stem cells)

L76 ANSWER 19 OF 35 USPATFULL on STN

ACCESSION NUMBER:

2003:240396 USPATFULL

TITLE:

Oligosaccharides, their preparation and pharmaceutical

compositions containing them

INVENTOR(S):

Mourier, Pierre, Charenton le Pont, FRANCE

Perrin, Elisabeth, Evrenus, FRANCE

Stutzmann, Jean-marie, Villecresnes, FRANCE Viskov, Christian, Ris Orangis, FRANCE

Wahl, Florence, Paris, FRANCE

PATENT ASSIGNEE (S):

Aventis Pharma, Cedex, FRANCE (non-U.S. corporation)

KIND NUMBER DATE _____

PATENT INFORMATION: APPLICATION INFO.:

US 6617316 B1 US 2000-693243 B1 20030909 20001020 (9)

NUMBER DATE

PRIORITY INFORMATION:

FR 1999-13182 19991022

US 2000-174647P 20000105 (60) Utility

DOCUMENT TYPE:

GRANTED

FILE SEGMENT: PRIMARY EXAMINER:

Fonda, Kathleen K.

LEGAL REPRESENTATIVE: NUMBER OF CLAIMS:

Newman, Irving

EXEMPLARY CLAIM:

35 1

NUMBER OF DRAWINGS:

0 Drawing Figure(s); 0 Drawing Page(s)

LINE COUNT:

798

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to oligosaccharides of formula: ##STR1##

to mixtures thereof, to diastereoisomers thereof, to a process for preparing them, to pharmaceutical compositions containing them, and to their use in preventing or treating a disease associated with an inflammatory process involving the production of nitric oxide.

9005-49-6, Reparin, reactions

(prepn. of uronic acid-contg. oligosaccharides as antiinflammatory

agents)

L76 ANSWER 20 OF 35 USPATFULL on STN

ACCESSION NUMBER:

2002:272935 USPATFULL

TITLE:

Novel differentiation inducing process of embryonic

stem cell to ectodermal cell and its use

INVENTOR(S):

Sasai, Yoshiki, Kyoto, JAPAN

Nishikawa, Shin-Ichi, Kyoto, JAPAN

10/644109

Page 27

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NUMBER
                                        KIND DATE
                           _____
                                      A1 20021017
A1 20010516 (9)
PATENT INFORMATION:
                       US 2002151056
                       US 2001-855587
APPLICATION INFO .:
                            NUMBER
                                         DATE
                       JP 2000-144059 20000516
PRIORITY INFORMATION:
                       JP 2000-290819
                                         20000925
                                         20001220 (60)
                       US 2000-257049P
                       Utility
DOCUMENT TYPE:
FILE SEGMENT:
                       APPLICATION
                       FITZPATRICK CELLA HARPER & SCINTO, 30 ROCKEFELLER
LEGAL REPRESENTATIVE:
                       PLAZA, NEW YORK, NY, 10112
NUMBER OF CLAIMS: .
EXEMPLARY CLAIM:
                       10 Drawing Page(s)
NUMBER OF DRAWINGS:
                       4056
LINE COUNT: ·
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
      A method for inducing differentiation of an embryonic stem cell into an
      ectodermal cell and an ectoderm-derived cell, which comprises culturing
      the embryonic stem cell under non-aggregation conditions; a medium and a
      medium supernatant used in the method; an agent for inducing
      differentiation used in the method; a stroma cell or a stroma
       cell-derived factor having activity of inducing differentiation in the
      method; an antibody which specifically recognizes the stroma cell; an
       antigen which recognizes the antibody; a cell induced by the method; a
      method for evaluating or screening a substance relating to the
       regulation in a differentiation step from an embryonic stem cell into an
       ectodermal cell or an ectoderm-derived cell by carrying out the method;
       and a medicament comprising the stroma cell, the stroma cell-derived
       cell, the antibody, the antigen or the cell.
     9005-49-6, Heparin, biological studies
IT
        (novel method for inducing differentiation of embryonic stem cells into
        ectodermal cells and use)
L76 ANSWER 21 OF 35 USPATFULL on STN
                       2002:72873 USPATFULL
ACCESSION NUMBER:
                       Novel therapeutic use of low molecular weight heparins
TITLE:
                       Stutzmann, Jean-Marie, Villecresnes, FRANCE
INVENTOR(S):
                       Uzan, Andre, Paris, FRANCE
                                       KIND DATE
                           NUMBER
                       _____
                       US 2002040013 A1 20020404 US 2001-881267 A1 20010614 (9)
PATENT INFORMATION:
APPLICATION INFO.:
                       Continuation of Ser. No. WO 1999-FR3109, filed on 13
RELATED APPLN. INFO.:
                       Dec 1999, UNKNOWN
                             NUMBER DATE
                       FR 1998-15919 19981217
PRIORITY INFORMATION:
                       Utility
DOCUMENT TYPE:
FILE SEGMENT:
                       APPLICATION
                       AVENTIS PHARMACEUTICALS, INC., PATENTS DEPARTMENT,
LEGAL REPRESENTATIVE:
                       ROUTE 202-206, P.O. BOX 6800, BRIDGEWATER, NJ,
                       08807-0800
                       19
NUMBER OF CLAIMS:
EXEMPLARY CLAIM:
                       1
LINE COUNT:
       The invention concerns the use of low molecular weight
       heparin for preventing and/or treating motor
```

neuron diseases.

L76 ANSWER 22 OF 35 USPATFULL on STN

ACCESSION NUMBER:

2002:32546 USPATFULL

TITLE:

Pharmaceutical compositions containing

oligosaccharides, the novel oligosaccharides and

preparation thereof

INVENTOR(S):

Mourier, Pierre, Charenton Le Pont, FRANCE

Perrin, Elisabeth, Evreux, FRANCE Viskov, Christian, Ris Orangis, FRANCE

NUMBER KIND DATE PATENT INFORMATION: US 2002019368 A1 20020214 US 6608042 B2 20030819 APPLICATION INFO.: US 2001-817428 A1 20010326 (9)

> DATE . NUMBER

PRIORITY INFORMATION:

FR 2000-3910 20000328 US 2000-205026P

Utility

20000518 (60)

DOCUMENT TYPE: FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

AVENTIS PHARMACEUTICALS, INC., PATENTS DEPARTMENT,

ROUTE 202-206, P.O. BOX 6800, BRIDGEWATER, NJ,

08807-0800

NUMBER OF CLAIMS:

36 1

EXEMPLARY CLAIM: LINE COUNT:

983

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to pharmaceutical compositions containing as an active ingredient at least one oligosaccharide of formula:

##STR1##

to novel oligosaccharides of formula (I), to mixtures thereof and to methods for their preparation.

ΙT 9005-49-6, Heparin, reactions

(prepn. of uronic acid-contg. oligosaccharides from heparin as antiinflammatory agents)

L76 ANSWER 23 OF 35 USPATFULL on STN

ACCESSION NUMBER:

2002:290736 USPATFULL

TITLE:

Identification of agents that protect against

inflammatory injury to neurons

INVENTOR(S):

Giulian, Dana, Houston, TX, United States

PATENT ASSIGNEE(S):

Baylor College of Medicine, Houston, TX, United States

(U.S. corporation)

NUMBER KIND DATE -----PATENT INFORMATION: US 6475745 B1 20021105 APPLICATION INFO.: US 1997-922889 19970903

RELATED APPLN. INFO.:

Division of Ser. No. US 1996-717551, filed on 20 Sep

DOCUMENT TYPE:

Utility GRANTED

FILE SEGMENT: PRIMARY EXAMINER:

Kunz, Gary L. Turner, Sharon

LEGAL REPRESENTATIVE:

Vinson & Elkins L.L.P.

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

ASSISTANT EXAMINER:

23

NUMBER OF DRAWINGS:

92 Drawing Figure(s); 38 Drawing Page(s)

LINE COUNT:

2755

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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Krishnan

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The present invention is directed to screening for an agent that inhibits the effect of a neurotoxin from a plaque component activated mononuclear phagocyte on a neuron. In addition, the present invention is directed to methods of screening for agents that inhibit mononuclear

directed to methods of screening for agents that inhibit mononuclear phagocyte-plaque component complex formation, plaque component activation of a mononuclear phagocyte, plaque component induced neurotoxicity of a mononuclear phagocyte. An agent obtained by the method of screening for an agent that inhibits mononuclear phagocyte-plaque component complex formation and a pharmaceutical

composition comprising the agent are embodied by the present invention. T 9005-49-6, Heparin sulfate, biological studies

(identification of agents that protect against inflammatory injury to neurons)

L76 ANSWER 24 OF 35 USPATFULL on STN

ACCESSION NUMBER:

2002:194555 USPATFULL

TITLE:

Composition of an endogenous insulin-like growth

factor-II derivative

INVENTOR(S):

Sakano, Katsuichi, Tokyo, JAPAN Higashihashi, Nobuyuki, Tokyo, JAPAN

Hashimoto, Ryuji, Tokyo, JAPAN

PATENT ASSIGNEE(S):

Daiichi Pharmaceutical Co., Ltd., Tokyo, JAPAN

(non-U.S. corporation)

NUMBER KIND DATE · B1 20020806 PATENT INFORMATION: US 6428781 19980709 WO 9829451 19990628 US 1999-331851 APPLICATION INFO.: WO 1997-JP4881 19971226 19990628 PCT 371 date

19990028 FCI 371 G

NUMBER DATE

PRIORITY INFORMATION:

JP 1996-349968 19961227

DOCUMENT TYPE:

Utility

FILE SEGMENT:

GRANTED

PRIMARY EXAMINER: ASSISTANT EXAMINER: Kemmerer, Elizabeth Bunner, Bridget E.

LEGAL REPRESENTATIVE:

Sughrue Mion, PLLC

NUMBER OF CLAIMS:

1

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

23 Drawing Figure(s); 12 Drawing Page(s)

LINE COUNT:

1427

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The effects of endogenous insulin-like growth factor can be appreciated by administering compounds capable of increasing free IGF in living bodies. Compounds are described which can elevate the concentration of unbound IGF by converting endogenous IGF (insulin-like growth factor) into free, biologically active IGF or elevating the concentration of the complex of IGF and IGFBP (insulin-like growth factor binding protein) in living bodies. Medicaments can be prepared containing these compounds or these compounds may be used in methods for the prevention and or treatment of IGF-responsive diseases such as diabetes, amyotrophic lateral sclerosis, or osteoporosis.

IT 9005-49-6, Heparin, biological studies

(IGF-II-IGFBP 3 binding inhibition by; methods and substances for elevating concn. of free insulin-like growth factor in vivo, and methods for screening substances for clin. use)

L76 ANSWER 25 OF 35 USPATFULL on STN

ACCESSION NUMBER:

2001:139285 USPATFULL

TITLE:

IDENTIFICATION OF AGENTS THAT PROTECT AGAINST

10/644109

Page 30

INFLAMMATORY INJURY TO NEURONS

INVENTOR(S): GIULIAN, DANA, HOUSTON, TX, United States

RELATED APPLN. INFO.: Division of Ser. No. US 1996-717551, filed on 20 Sep

1996, GRANTED, Pat. No. US 6071493

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: VINSON & ELKINS L.L.P., ATTN: DOCKET SPECIALIST, 2300

FIRST CITY TOWER, HOUSTONLPHIA, TX, 770026760

NUMBER OF CLAIMS: 9 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 29 Drawing Page(s)

LINE COUNT: 2790

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention is directed to screening for an agent that inhibits the effect of a neurotoxin from a plaque component activated mononuclear phagocyte on a neuron. In addition, the present invention is directed to methods of screening for agents that inhibit mononuclear phagocyte-plaque component complex formation, plaque component activation of a mononuclear phagocyte, plaque component induced neurotoxicity of a mononuclear phagocyte. An agent obtained by the method of screening for an agent that inhibits mononuclear phagocyte-plaque component complex formation and a pharmaceutical composition comprising the agent are embodied by the present invention.

IT 9005-49-6, Heparin sulfate, biological studies

L76 ANSWER 26 OF 35 USPATFULL on STN

ACCESSION NUMBER: 2001:139284 USPATFULL

TITLE; IDENTIFICATION OF AGENTS THAT PROTECT AGAINST

INFLAMMATORY INJURY TO NEURONS

INVENTOR(S): GIULIAN, DANA, HOUSTON, TX, United States

RELATED APPLN. INFO.: Division of Ser. No. US 1996-717551, filed on 20 Sep

1996, GRANTED, Pat. No. US 6071493

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: VINCE & ELKINS, L.L.P., ATTN: DOCKET SPECIALIST, 2300

FIRST CITY TOWER, 1001 FANNIN STREET, HOUSTON, TX,

770026760

NUMBER OF CLAIMS: 99
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 29 Drawing Page(s)

LINE COUNT: 2792

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention is directed to screening for an agent that inhibits the effect of a neurotoxin from a plaque component activated mononuclear phagocyte on a neuron. In addition, the present invention is directed to methods of screening for agents that inhibit mononuclear phagocyte-plaque component complex formation, plaque component activation of a mononuclear phagocyte, plaque component induced neurotoxicity of a mononuclear phagocyte. An agent obtained by the

method of screening for an agent that inhibits mononuclear phagocyte-plaque component complex formation and a pharmaceutical composition comprising the agent are embodied by the present invention.

9005-49-6, Heparin sulfate, biological studies (identification of agents that protect against inflammatory injury to

L76 ANSWER 27 OF 35 USPATFULL on STN

ACCESSION NUMBER:

2000:70424 USPATFULL

TITLE:

ΙT

Method of screening for an agent that inhibits mononuclear phagocyte-plaque component complex

formation

INVENTOR (\$):

Giulian, Dana, Houston, TX, United States

PATENT ASSIGNEE(S):

Baylor College of Medicine, Houston, TX, United States

DATE

(U.S. corporation)

KIND NUMBER US 6071493 · 20000606 PATENT INFORMATION: US 1996-717551 19960920 APPLICATION INFO .:

DOCUMENT TYPE:

Utility Granted

FILE SEGMENT:

Duffy, Patricia A. PRIMARY EXAMINER: Corder, Timothy S. Vinson & Elkins LLP LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS:

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

92 Drawing Figure(s); 38 Drawing Page(s)

2660

LINE COUNT: CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention is directed to screening for an agent that inhibits mononuclear phagocyte-plaque component complex formation (hereinafter "complex formation"). The methods include the steps of contacting a mononuclear phagocyte with a plaque component to stimulate complex formation and adding an agent suspected of inhibiting complex formation, measuring complex formation, and comparing complex formation to a measured control, wherein the reduction of complex formation compared to the control results in detection of an agent that inhibits complex formation. The mononuclear phagocytes may be from mammalian brain. The plaque component may be coupled to a solid support.

9005-49-6, Heparin sulfate, biological studies ΙT

(identification of agents that protect against inflammatory injury to neurons)

L76 ANSWER 28 OF 35 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: DOCUMENT NUMBER:

1997:308923 BIOSIS PREV199799616726

TITLE:

Effects of low doses of glycosaminoglycans and insulin-like

growth factor-I on motor neuron

disease in wobbler mouse.

AUTHOR (S):

Vergani, Letizia; Finco, Cristina; Di Giulio, Anna Maria;

Muller, E. E.; Gorio, Alfredo [Reprint author]

CORPORATE SOURCE:

Lab. Res. Pharmacol. Neurodegenerative Disorders, Dep. Pharmacol. Chemotherapy Med. Toxicol., Univ. Milano, Via

SOURCE:

Vanvitelli 32, 20124 Milano, Italy Neuroscience Letters, (1997) Vol. 228, No. 1, pp. 41-44.

CODEN: NELED5. ISSN: 0304-3940.

DOCUMENT TYPE:

Article

LANGUAGE: ENTRY DATE: English Entered STN: 26 Jul 1997

Last Updated on STN: 26 Jul 1997

```
ABSTRACT:In this study we examined the effects of insulin-like growth factor-I (IGF-I) and of glycosaminoglycans (GAGs) on the progressive motor
***neuron*** disease in wobbler mice. After clinical diagnosis at
age 3 weeks, mice received daily subcutaneous injections of IGF-I, or GAGs, or
saline for 3 weeks. The histometric analysis revealed that biceps muscle fiber
diameter was reduced in wobbler mice and that treatments with GAGs and IGF-I
provented such a drop. The number of atrophic small fibers was markedly
reduced and that of the larger ones augmented. No effects on body growth and
biceps muscle weight were observed. The combined AChE-silver staining revealed
that both treatments promoted intramuscular axonal sprouting. The typical
decline of grip strength in wobbler mice was also prevented. This study
suggests that GAGs and IGF-I administrations can retard the onset of motor
deficit, and reduce muscle atrophy in wobbler mice.
```

CONCEPT CODE:

Biochemistry studies - Proteins, peptides and amino acids

10064

Biochemistry studies - Carbohydrates 10068

Pathology - Necrosis 12510 Pathology - Therapy 12512 Endocrine - General 17002

Endocrine - Neuroendocrinology 17020

Muscle - Physiology and biochemistry 17504

Muscle - Pathology 17506

Integumentary system - General and methods 18501 Nervous system - Physiology and biochemistry 20504

Nervous system - Pathology 20506
Pharmacology - Endocrine system 22016
Pharmacology - Muscle system 22022
Pharmacology - Neuropharmacology 22024

Routes of immunization, infection and therapy 22100 Development and Embryology - Morphogenesis 25508

Laboratory animals - General 28002

INDEX TERMS:

Major Concepts
Animal Care; Biochemistry and Molecular Biophysics;

Development; Endocrine System (Chemical Coordination and

Homeostasis); Integumentary System (Chemical

Coordination and Homeostasis); Muscular System (Movement

and Support); Nervous System (Neural Coordination);

Pharmacology

INDEX TERMS: Chemicals & Biochemicals

INSULIN-LIKE GROWTH FACTOR-I; HEPARIN

INDEX TERMS:

Miscellaneous Descriptors

ADMINISTRATION; ANIMAL MODEL; BICEPS MUSCLE; GLYCOSAMINOGLYCANS; HEPARIN; INSULIN-LIKE GROWTH FACTOR-I; INTRAMUSCULAR AXONAL SPROUTING; LOW DOSES;

MOTOR DEFICIT; MOTOR NEURON DEGENERATION; MOTOR

NEURON DISEASE; MUSCLE ATROPHY; MUSCLE

DISEASE; MUSCLE STRENGTH; MUSCLE WEIGHT; MUSCULAR SYSTEM; NERVE ATROPHY; NERVOUS SYSTEM; NERVOUS SYSTEM DISEASE; PHARMACOLOGY; SUBCUTANEOUS INJECTION; WOBBLER

MOUSE

ORGANISM:

Classifier

Muridae · 86375

Super Taxa

Rodentia; Mammalia; Vertebrata; Chordata; Animalia

Organism Name Muridae Taxa Notes

Animals, Chordates, Mammals, Nonhuman Vertebrates,

Nonhuman Mammals, Rodents, Vertebrates

REGISTRY NUMBER:

67763-96-6 (INSULIN-LIKE GROWTH FACTOR-I)

9005-49-6 (HEPARIN)

L76 ANSWER 29 OF 35 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN

10/644109

Page 33

ACCESSION NUMBER:

1986:120194 BIOSIS

DOCUMENT NUMBER:

PREV198681030610; BA81:30610

TITLE:

MULTIPLE RETINAL MUSCULAR AND CUTANEOUS CHOLESTEROL EMBOLI

A CASE WITH PROGRESSIVE ENCEPHALOPATHY.

AUTHOR(S):

BUGE A [Reprint author]; VINCENT D; RANCUREL G; BAUDRIMONT

M; DUBAS F; HAUW J-J

CORPORATE SOURCE:

CLIN NEUROL, HOPITAL DE LA SALPETRIERE, 47 BLVD

DEL'HOPITAL, F-75651 PARIS CEDEX 13, FR

SOURCE:

Revue Neurologique (Paris), (1985) Vol. 141, No. 8-9, pp.

578-582.

CODEN: RENEAM. ISSN: 0035-3787.

DOCUMENT TYPE:

Article

FILE SEGMENT:

FRENCH

LANGUAGE: ENTRY DATE:

Entered STN: 25 Apr 1986

Last Updated on STN: 25 Apr 1986

ABSTRACT: A 73 year-old man experienced left monocular blindness and transient right hand clumsiness. A left carotid arteriogram was performed 4 days after admission. Immediately following arteriography, there was a right hemiparesia and dysphasia. After 24 hours, the abnormalities resolved. The patient was treated with heparin. During the following weeks, he became gradually drowsy and confused. Pseudo-bulbar palsy and astasia appeared after a fluctuating but progressive neurological course. The combination of systemic symptoms, high sedimentation rate, renal failure, livedo reticularis and purple toes suggested necrotizing angiitis. With corticosteroid treatment, there was a slight improvement of systemic symptoms. Cholesterol emboli were seen in both fundi. Cholesterol embolization was proved by identifying the biconcave cholesterol crystal clefts in muscle and skin biopsies. The subsequent course was marqued by continuous neurological deterioration. patient became stuporous and died 7 months after admission. Despite the lack of central nervous system pathological study, the clinical picture was highly suggestive of cerebral cholesterol embolism. A few cases have been reported, with only eight well-documented clinical descriptions. Clinical signs and symptoms were closely similar to those of the present case. Anticoagulant therapy of cholesterol emboli has been unsuccessful. In the present case the onset of embolization was temporally related to anticoagulation. CONCEPT CODE:

Biochemistry studies - Sterols and steroids

Biochemistry studies - Carbohydrates

Anatomy and Histology - Experimental anatomy

Pathology - Necrosis 12510

Metabolism - Sterols and steroids

Cardiovascular system - Blood vessel pathology

Endocrine - Adrenals 17004 17506 Muscle - Pathology

Integumentary system - Pathology 18506

Sense organs - Pathology 20006 Nervous system - Pathology

Pharmacology - Clinical pharmacology 22005 Pharmacology - Cardiovascular system

Pharmacology - Endocrine system 22016

INDEX TERMS:

Major Concepts

Cardiovascular Medicine (Human Medicine, Medical Sciences); Dcrmatology (Human Medicine, Medical

Sciences); Endocrine System (Chemical Coordination and Homeostasis); Metabolism; Muscular System (Movement and Support); Neurology (Human Medicine, Medical Sciences);

Pharmacology; Sense Organs (Sensory Reception)

INDEX TERMS:

Miscellaneous Descriptors

HUMAN CORTICOSTEROID HORMONE-DRUG HEPARIN

CARDIOVASCULAR-DRUG ANTICOAGULANT-DRUG DYSPHASIA

HEMIPARESIS NECROTIZING ANGIITIS

ORGANISM:

Classifier

Hominidae 86215

BEST AVAILABLE COPY

Krishnan

10/644109

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Super Taxa

Primates; Mammalia; Vertebrata; Chordata; Animalia

Taxa Notes

Animals, Chordates, Humans, Mammals, Primates,

Vertebrates

REGISTRY NUMBER:

57-88-5 (CHOLESTEROL) 9005-49-6 (HEPARIN)

L76 ANSWER 30 OF 35 EMBASE COPYRIGHT 2003 ELSEVIER INC. ALL RIGHTS RESERVED.

on STN

ACCESSION NUMBER:

2003247308 EMBASE

TITLE:

Strokes, thromboembolic events, and IVIg: Rare incidents

blemish an excellent safety record.

AUTHOR:

Dalakas M.C.; Clark W.M.

CORPORATE SOURCE:

Dr. M.C. Dalakas, Neuromuscular Diseases Section, NINDS, NIH, 10 Center Dr., Bethesda, MD 20892-1382, United States.

dalakasm@ninds.nih.gov

SOURCE:

Neurology, (10 Jun 2003) 60/11 (1736-1737).

Refs: 10

ISSN: 0028-3878 CODEN: NEURAI

COUNTRY:

United States

DOCUMENT TYPE:

Journal; Editorial

FILE SEGMENT:

Neurology and Neurosurgery

026 Immunology, Serology and Transplantation

037 Drug Literature Index

038 Adverse Reactions Titles

LANGUAGE:

English

CONTROLLED TERM: ·

Medical Descriptors:

*stroke: DT, drug therapy *stroke: PC, prevention *stroke: SI, side effect

*occlusive cerebrovascular disease: DT, drug therapy *occlusive cerebrovascular disease: PC, prevention *occlusive cerebrovascular disease: SI, side effect

thromboembolism: DT, drug therapy thromboembolism: PC, prevention thromboembolism: SI, side effect autoimmune disease: DT, drug therapy neuromuscular disease: DT, drug therapy

drug efficacy drug safety

motor neuron disease: DT, drug therapy Guillain Barre syndrome: DT, drug therapy

chronic inflammatory demyelinating polyneuropathy: DT, drug

therapy

myasthenia: DT, drug therapy dermatomyositis: DT, drug therapy

Eaton Lambert syndrome: DT, drug therapy stiff man syndrome: DT, drug therapy.

drug mechanism

complement activation

macrophage

side effect: SI, side effect headache: SI, side effect chill: SI, side effect myalgia: SI, side effect low back pain: SI, side effect thorax pain: SI, side effect

aseptic meningitis: SI, side effect

rash: SI, side effect

anaphylaxis: SI, side effect immunoglobulin A deficiency

```
kidney tubule necrosis: SI, side effect
kidney disease
high risk patient
vascular disease
treatment outcome
fibrinolytic therapy
disease predisposition
muscle weakness
immobilization
wheelchair
lung embolism: SI, side effect
drug infusion
antibody blood level
capillary flow
hypergammaglobulinemia
hypercholesterolemia
risk factor
blood flow velocity
hydration
thrombocytosis
diabetes mellitus
prophylaxis
echography
low drug dose
human
clinical trial
editorial
priority journal
Drug Descriptors:
*immunoglobulin: AE, adverse drug reaction
*immunoglobulin: CT, clinical trial
*immunoglobulin: CB, drug combination
*immunoglobulin: DT, drug therapy
*immunoglobulin: PD, pharmacology
*immunoglobulin: IV, intravenous drug administration
autoantibody: EC, endogenous compound
complement: EC, endogenous compound
cytokine: EC, endogenous compound
Fc receptor: EC, endogenous compound
immunoglobulin A: EC, endogenous compound
tissue plasminogen activator: DT, drug therapy
tissue plasminogen activator: PD, pharmacology
placebo
immunoglobulin G: EC, endogenous compound
fibrinogen: EC, endogenous compound
phospholipid antibody: EC, endogenous compound
acetylsalicylic acid: DT, drug therapy
acetylsalicylic acid: PD, pharmacology
nimodipine: DT, drug therapy
nimodipine: PD, pharmacology
anticoagulant agent: DT, drug therapy
anticoagulant agent: PD, pharmacology
urokinase: DT, drug therapy
urokinase: PD, pharmacology
heparin: DT, drug therapy
heparin: PD, pharmacology
  low molecular weight heparin: CB, drug combination
  low molecular weight heparin: DT, drug therapy
  low molecular weight heparin: PD, pharmacology
(immunoglobulin) 9007-83-4; (complement) 9007-36-7; (tissue
plasminogen activator) 105913-11-9; (immunoglobulin G)
97794-27-9; (fibrinogen) 9001-32-5; (acetylsalicylic acid)
493-53-8, 50-78-2, 53663-74-4, 53664-49-6, 63781-77-1;
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CAS REGISTRY NO.:

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(nimodipine) 66085-59-4; (urokinase) 139639-24-0; (heparin)

37187-54-5, 8057-48-5, 8065-01-8, 9005-48-5

CHEMICAL NAME: Aspirin

L76 ANSWER 31 OF 35 EMBASE COPYRIGHT 2003 ELSEVIER INC. ALL RIGHTS RESERVED.

on STN

ACCESSION NUMBER: 2002190914 EMBASE

TITLE: Caesarean section

Caesarean section conducted under subarachnoid block in two

sisters with spinal muscular atrophy.

AUTHOR: Harris S.J.; Moaz K.

CORPORATE SOURCE: S.J. Harris, Queen Elizabeth Hospital, Gayton Road,

Norfolk, PE30 4ET, United Kingdom. angela.kent@klshosp.anglox.nhs.uk

SOURCE: International Journal of Obstetric Anesthesia, (2002) 11/2

(125-127). Refs: 10

ISSN: 0959-289X CODEN: IOANER

COUNTRY: United Kingdom
DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 024 Anesthesiology

037 Drug Literature Index

LANGUAGE: English
SUMMARY LANGUAGE: English

ABSTRACT:

Spinal muscular atrophy is a rare chronic neurological condition characterised by degeneration of the anterior horn cell. Experience with the anaesthetic management of the pregnant patient with this condition is limited. We report the clinical details of two wheelchair-bound sisters, who underwent elective caesarean section within a few weeks of one another. Both patients were safely managed with subarachnoid anaesthesia without any deterioration of their underlying neurological condition. It is hoped that this report will add to the evidence that subarachnoid anaesthesia can safely be used for caesarean section in chronic neurological conditions and, in particular, spinal muscular atrophy. COPYRGT. 2002 published by Elsevier Science Ltd.

CONTROLLED TERM: Medical Descriptors:

*spinal muscular atrophy

*cesarean section
*obstetric anesthesia
neurologic disease
elective surgery
general anesthesia
regional anesthesia

postoperative pain: CO, complication postoperative pain: DT, drug therapy postoperative pain: PC, prevention

subarachnoid space chronic disease

thromboembolism: CO, complication thromboembolism: DT, drug therapy thromboembolism: PC, prevention

human female case report adult article

Drug Descriptors:

*bupivacaine: AD, drug administration

*bupivacaine: DO, drug dose

*bupivacaine: EI, epidural drug administration

morphine: AD, drug administration

morphine: DT, drug therapy

morphine: EI, epidural drug administration

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morphine: IM, intramuscular drug administration

diclofenac: AD, drug administration

diclofenac: DT, drug therapy

diclofenac: RC, rectal drug administration low molecular weight heparin: DO, drug dose low molecular weight heparin: DT, drug therapy low molecular weight heparin: SC, subcutaneous drug

administration

enoxaparin: DO, drug dose enoxaparin: DT, drug therapy

enoxaparin: SC, subcutaneous drug administration

fentanyl: AD, drug administration
fentanyl: DO, drug dose

fentanyl: EI, epidural drug administration anesthetic agent: AD, drug administration

anesthetic agent: DO, drug dose.

anesthetic agent: EI, epidural drug administration

suxamethonium rocuronium

neuromuscular blocking agent

(bupivacaine) 18010-40-7, 2180-92-9, 55750-21-5; (morphine) CAS REGISTRY NO.:

52-26-6, 57-27-2; (diclofenac) 15307-79-6, 15307-86-5;

(enoxaparin) 9041-08-1; (fentanyl) 437-38-7;

(suxamethonium) 306-40-1, 71-27-2; (rocuronium) 119302-91-9

CHEMICAL NAME: Clexane

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· on STN

2001277576 EMBASE

ACCESSION NUMBER: TITLE:

Hodgkin's disease complicated by the nephrotic syndrome in

a man with Kugelberg-Welander disease.

AUTHOR: Thomson J.A.; Seymour J.F.; Wolf M.

M. Wolf, Div. of Haematology/Medical Oncology, Peter CORPORATE SOURCE:

MacCallum Cancer Institute, A'Beckett Street, Melbourne,

Vic. 8006, Australia

Leukemia and Lymphoma, (2001) 42/3 (561-566). SOURCE:

Refs: 23

ISSN: 1042-8194 CODEN: LELYEA

COUNTRY:

United Kingdom Journal; Article

DOCUMENT TYPE: FILE SEGMENT:

General Pathology and Pathological Anatomy 005

016 Cancer Hematology 025

Urology and Nephrology 028 037 Drug Literature Index

LANGUAGE:

English English

SUMMARY LANGUAGE:

A case of nephrotic syndrome due to minimal change glomerulonephritis complicating Hodgkin's disease in a man with a longstanding neurological disorder is presented. Treatment with combination chemotherapy resulted in a rapid improvement in the nephrotic syndrome, and complete remission of the Hodgkin's disease. Disease relapse occurred less than 12 months later without recurrence of the nephrotic syndrome and was refractory to further treatment. The association of minimal change glomerulonephritis with Hodgkin's disease and the possible pathogenesis of this association are discussed.

CONTROLLED TERM: Medical Descriptors:

> *Hodgkin disease: DI, diagnosis *Hodgkin disease: DT, drug therapy *nephrotic syndrome: CO, complication *nephrotic syndrome: DI, diagnosis *nephrotic syndrome: DT, drug therapy

```
*nephrotic syndrome: ET, etiology
                      *Kugelberg Welander disease: DI, diagnosis
                    complication: CO, complication
                    minimal change glomerulonephritis: DI, diagnosis
                    neurologic disease
                    cancer combination chemotherapy
                    drug response
                    leukemia remission
                    cancer recurrence
                    recurrent disease
                    disease association
                    pathogenesis
                    kidney biopsy
                    laboratory diagnosis
                    human
                    male
                    case report
                    controlled study
                    human tissue
                    adult
                    article
                    priority journal
                    Drug Descriptors:
                      enoxaparin: DT, drug therapy
                      enoxaparin: SC, subcutaneous drug administration
                    dexamethasone: DT, drug therapy
                    allopurinol: DT, drug therapy
                    prednisolone: CB, drug combination
                    prednisolone: DT, drug therapy
                    cyclophosphamide: CB, drug combination
                    cyclophosphamide: DT, drug therapy
                    etoposide: CB, drug combination
                    etoposide: DT, drug therapy
                    procarbazine: CB, drug combination
procarbazine: DT, drug therapy
                    doxorubicin: CB, drug combination
                    doxorubicin: DT, drug therapy
                    bleomycin: CB, drug combination
                    bleomycin: DT, drug therapy
                    (enoxaparin) 9041-08-1; (dexamethasone) 50-02-2;
                    (allopurinol) 315-30-0; (prednisolone) 50-24-8;
                    (cyclophosphamide) 50-18-0; (etoposide) 33419-42-0;
                    (procarbazine) 366-70-1, 671-16-9; (doxorubicin)
                    23214-92-8, 25316-40-9; (bleomycin) 11056-06-7
L76 ANSWER 33 OF 35 EMBASE COPYRIGHT 2003 ELSEVIER INC. ALL RIGHTS RESERVED.
                    1999203917 EMBASE
                    [Report from the USA].
                    BERICHT AUS USA.
                    Gakenheimer W.C.
                    Dr. W.C. Gakenheimer, 413 Stafford Road, Wilmington, DE
                    19803, United States
                    Pharmazeutische Industrie, (1999) 61/5 (449-452).
                    ISSN: 0031-711X CODEN: PHINAN
                    Germany
                    Journal; Article
```

CAS REGISTRY NO.:

on STN

TITLE:

AUTHOR:

SOURCE:

COUNTRY:

LANGUAGE:

ACCESSION NUMBER:

CORPORATE SOURCE:

DOCUMENT TYPE:

FILE SEGMENT:

Medical Descriptors: CONTROLLED TERM: *drug packaging

037

039

German

Drug Literature Index

Pharmacy

```
*drug industry
```

food and drug administration

standardization

amyotrophic lateral sclerosis

multiple sclerosis: DT, drug therapy

unstable angina pectoris: DT, drug therapy

oncogene neu

prostate cancer: DT, drug therapy depression: DT, drug therapy

article

Drug Descriptors: .

mevinolin fluoxetine

protein c: EC, endogenous compound

ligand glatiramer

dalteparin: DT, drug therapy

prostate specific antigen: EC, endogenous compound

taxol

doxorubicin

betala interferon: DT, drug therapy

paroxetine

venlafaxine: DT, drug therapy

antineoplastic agent: DT, drug therapy

mdx 210

(mevinolin) 75330-75-5; (fluoxetine) 54910-89-3, CAS REGISTRY NO.:

56296-78-7, 59333-67-4; (protein c) 60202-16-6;

(glatiramer) 147245-92-9, 28704-27-0; (taxol) 33069-62-4;

(doxorubicin) 23214-92-8, 25316-40-9; (paroxetine)

61869-08-7; (venlafaxine) 93413-69-5

(1) Copaxone; (2) Fragmin; (3) Mdx 210; (4) Paxil; (5) CHEMICAL NAME:

Effexor; (6) Rebif; (7) Mevacor; (8) Prozac

(1) Teva (Israel); (2) Pharmacia Upjohn (United States); COMPANY NAME:

(3) Medarex (United States); (4) Smith Kline Beecham; (5) Wyeth Ayerst; (6) Serono; (7) Merck; (8) Lilly; Neopharma

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96288534 EMBASE ACCESSION NUMBER:

DOCUMENT NUMBER:

1996288534

TITLE:

Clinical pharmacology and therapeutics.

Kochar M.S.; Campbell W.B. AUTHOR:

Department of Medicine, Zablocki VA Medical Center, Medical CORPORATE SOURCE:

College of Wisconsin, Milwaukee, WI, United States Wisconsin Medical Journal, (1996) 95/9 (645-646).

ISSN: 0043-6542 CODEN: WMJOA7

COUNTRY:

SOURCE:

United States

DOCUMENT TYPE: FILE SEGMENT:

Journal; (Short Survey) Internal Medicine 006

030 Pharmacology

Drug Literature Index 037 Adverse Reactions Titles 038

LANGUAGE:

English

CONTROLLED TERM:

Medical Descriptors: *clinical pharmacology

abortion

amyotrophic lateral sclerosis: DT, drug therapy

chickenpox: DT, drug therapy chickenpox: PC, prevention

gastrointestinal toxicity: SI, side effect

general anesthesia

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```
headache: SI, side effect
hepatitis a: PC, prevention
hepatitis a: DT, drug therapy
human
human immunodeficiency virus infection: DT, drug therapy
hypertension: DT, drug therapy
intramuscular drug administration
intravaginal drug administration
lactic acidosis: SI, side effect
migraine: DT, drug therapy
non insulin dependent diabetes mellitus: DT, drug therapy
obesity: DT, drug therapy
oral drug administration
osteoporosis: DT, drug therapy
short survey
subcutaneous drug administration
Drug Descriptors:
*alendronic acid: DT, drug therapy
*alendronic acid: PD, pharmacology
*anti human immunodeficiency virus agent: DT, drug therapy
*anti human immunodeficiency virus agent: CB, drug
combination
*anti human immunodeficiency virus agent: PD, pharmacology
*antidiabetic agent: AE, adverse drug reaction
*antidiabetic agent: DT, drug therapy
*antidiabetic agent: PD, pharmacology
*dexfenfluramine: PD, pharmacology
*dexfenfluramine: DT, drug therapy
*dipeptidyl carboxypeptidase inhibitor: CM, drug comparison
*dipeptidyl carboxypeptidase inhibitor: DT, drug therapy
*dipeptidyl carboxypeptidase inhibitor: PD, pharmacology
*losartan potassium: CM, drug comparison
*losartan potassium: DT, drug therapy
*losartan potassium: PD, pharmacology
abortive agent: AE, adverse drug reaction abortive agent: PD, pharmacology
acarbose: AE, adverse drug reaction
acarbose: PD, pharmacology
acarbose: DT, drug therapy
angiotensin receptor antagonist: CM, drug comparison
angiotensin receptor antagonist: DT, drug therapy
angiotensin receptor antagonist: PD, pharmacology
biguanide: PD, pharmacology
biguanide: DT, drug therapy
biguanide: AE, adverse drug reaction
biguanide: CB, drug combination
bisphosphonic acid derivative: PD, pharmacology
bisphosphonic acid derivative: DT, drug therapy
chickenpox vaccine: DT, drug therapy
  dalteparin: DT, drug therapy
desflurane
  enoxaparin: DT, drug therapy
hepatitis a vaccine: DT, drug therapy
indinavir: CB, drug combination
indinavir: DT, drug therapy
indinavir: PD, pharmacology
inhalation anesthetic agent
  low molecular weight heparin: DT, drug therapy
metformin: PD, pharmacology
metformin: AE, adverse drug reaction
metformin: DT, drug therapy
metformin: CB, drug combination
methotrexate: AE, adverse drug reaction
```

CAS REGISTRY NO .:

CHEMICAL NAME:

DOC. NO. CPI:

DERWENT CLASS:

COUNTRY COUNT:

PATENT NO

FR 2787329

INVENTOR(S):

TITLE:

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```
methotrexate: PD, pharmacology
                   misoprostol: PD, pharmacology
                   misoprostol: AE, adverse drug reaction
                   nucleoside derivative: DT, drug therapy
                   nucleoside derivative: CB, drug combination
                   proteinase inhibitor: CB, drug combination
                   proteinase inhibitor: DT, drug therapy
                   proteinase inhibitor: PD, pharmacology
                   riluzole: DT, drug therapy
                   riluzole: PD, pharmacology
                   ritonavir: DT, drug therapy
                   ritonavir: PD, pharmacology
                    ritonavir: CB, drug combination
                    saquinavir: PD, pharmacology
                    saquinavir: CB, drug combination
                    saquinavir: DT, drug therapy
                    sevoflurane
                    sulfonylurea: DT, drug therapy
                    sulfonylurea: CB, drug combination
                    sumatriptan: DT, drug therapy
                    sumatriptan: PD, pharmacology
                    sumatriptan succinate
                    (alendronic acid) 66376-36-1; (dexfenfluramine) 3239-44-9,
                   3239-45-0; (losartan potassium) 124750-99-8; (acarbose)
                    56180-94-0; (biguanide) 56-03-1; (desflurane) 57041-67-5;
                    (enoxaparin) 9041-08-1; (indinavir) 150378-17-9,
                    157810-81-6; (metformin) 1115-70-4, 657-24-9;
                    (methotrexate) 15475-56-6, 59-05-2, 7413-34-5;
                    (misoprostol) 59122-46-2, 59122-48-4; (proteinase
                    inhibitor) 37205-61-1; (riluzole) 1744-22-5; (ritonavir) .
                    155213-67-5; (saquinavir) 127779-20-8; (sevoflurane)
                    28523-86-6; (sumatriptan) 103628-46-2; (sumatriptan
                    succinate) 103628-48-4
                   Cozaar; Invirase; Norvir; Crixivan; Glucophage; Precose;
                    Fosamax; Redux; Imitrex; Rilutek; Folex; Cytotec; Lovenox;
                    Fragmin; Varivax; Havrix; Suprane
L76 ANSWER 35 OF 35 WPIDS COPYRIGHT 2003 THOMSON DERWENT on STN
                      2000-442268 [38] WPIDS
ACCESSION NUMBER:
                      C2000-134436
                      Use of low molecular weight heparin
                      for treatment and prevention of motor
                      neuron disease, e.g. amyotrophic
                      lateral sclerosis.
                      B04 ·
                   STUTZMANN, J M; UZAN, A; STUTZMANN, J
                      (AVET) AVENTIS PHARMA SA; (STUT-1) STUTZMANN J; (UZAN-I)
PATENT ASSIGNEE(S):
                     ·UZAN A
                     . 83
PATENT INFORMATION:
               KIND DATE
                            WEEK
                                         LA
                                              PG MAIN IPC
     WO 2000035462 A1 20000622 (200038)* FR 18 A61K031-727
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL
           OA PT SD SE SL SZ TZ UG ZW
         W: AE AL AU BA BB BG BR CA CN CR CU CZ DM EE GD GE HR HU ID IL IN IS
            JP KP KR LC LK LR LT LV MA MG MK MN MX NO NZ PL RO RU SG SI SK SL
            TR TT UA US UZ VN YU ZA
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AU 2000015697 A 20000703 (200046) A61K031-727 NO 2001002849 A 20010608 (200154) A61K000-00 A1 20011010 (200167) FR A61K031-727 EP 1140119

A1 20000623 (200038)

A61K031-738

10/644109

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1...

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI

US 2002040013 A1 20020404 (200227)

A61K031-727 19 A61K031-727

.JP 2002532431 W 20021002 (200279)

APPLICATION DETAILS:

PATENT NO KIND	APPLICATION	DATE
WO 2000035462 A1	WO 1999-FR3109	19991213
FR 2787329 A1	FR 1998-15919	19981217
AU 2000015697 A	AU 2000-15697	19991213
NO 2001002849 A	WO 1999-FR3109	19991213
	· NO 2001-2849	20010608
EP 1140119 A1	EP 1999-958308	19991213
. ·	WO 1999-FR3109	19991213
US 2002040013 A1 Cont of	WO 1999-FR3109	19991213
	US 2001-881267	20010614
JP 2002532431 W	WO 1999-FR3109	19991213
	JP 2000-587782	19991213

FILING DETAILS:

PATENT NO K		PATENT NO
AU 2000015697 EP 1140119 JP 2002532431	A Based on Al Based on	WO 2000035462 WO 2000035462 WO 2000035462

PRIORITY APPLN. INFO: FR 1998-15919 19981217

INT. PATENT CLASSIF.:

MAIN:

A61K000-00; A61K031-727; A61K031-738.

SECONDARY:

A61P009-10; A61P025-00; A61P025-28; A61P043-00

ADDITIONAL:

C08B037-10

BASIC ABSTRACT:

WO 200035462 A UPAB: 20000811

NOVELTY - Use of low molecular weight heparin (I) to

produce a medicine that promotes survival and/or growth of motor neurons.

ACTIVITY - Cytoprotective; neurotrophic.

A mixed culture of astrocytes and motor neurons (MN) was treated with the low molecular weight heparin Enoxaparine

(Ia), then after 2-3 days the number of viable MN assessed from:

(i) immunoreactivity for the homoprotein Islet1/2 or for neurofilaments; and

(ii) presence of neurites longer than 10 cell diameters.

At 10 ng/ml (Ia), the mean number of MN was 196% and the mean MN survival was 120.7%, both relative to a vehicle-only control as 100%. The number of very large MN was 66 per cubic centimeters (cc) in presence of (Ia) compared with 38 per cc in a control.

MECHANISM OF ACTION - None given.

No biological data given.

USE - (I) is specifically used to treat and/or prevent motor neuron diseases, particularly amyotrophic lateral sclerosis, progressive spinal muscular atrophy and infantile muscular atrophy.

Dwg.0/0

FILE SEGMENT: CPI FIELD AVAILABILITY: AB; DCN

MANUAL CODES:

CPI: B04-C02E; B14-J01; B14-J05A